



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification ⁷ : C07K 14/00	A2	(11) International Publication Number: WO 00/61612 (43) International Publication Date: 19 October 2000 (19.10.00)
(21) International Application Number: PCT/US00/08896 (22) International Filing Date: 3 April 2000 (03.04.00) (30) Priority Data: 09/285,479 2 April 1999 (02.04.99) US 09/466,396 17 December 1999 (17.12.99) US 09/476,496 30 December 1999 (30.12.99) US 09/480,884 10 January 2000 (10.01.00) US 09/510,376 22 February 2000 (22.02.00) US (71) Applicant (for all designated States except US): CORIXA CORPORATION [US/US]; Suite 200, 1124 Columbia Street, Seattle, WA 98104 (US). (72) Inventors; and (75) Inventors/Applicants (for US only): WANG, Tongtong [US/US]; 8049 NE 28th Street, Medina, WA 98039 (US). FAN, Lijun [CN/US]; 14116 SE 46th Street, Bellevue, WA 98006 (US). (74) Agents: MAKI, David, J.; Seed Intellectual Property Law Group PLLC, Suite 6300, 701 Fifth Avenue, Seattle, WA 98104-7092 (US) et al.		(81) Designated States: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PE, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG). Published <i>Without international search report and to be republished upon receipt of that report.</i>
(54) Title: COMPOUNDS AND METHODS FOR THERAPY AND DIAGNOSIS OF LUNG CANCER (57) Abstract <p>Compounds and methods for the treatment and diagnosis of lung cancer are provided. The inventive compounds include polypeptides containing at least a portion of a lung tumor protein. Vaccines and pharmaceutical compositions for immunotherapy of lung cancer comprising such polypeptides, or DNA molecules encoding such polypeptides, are also provided, together with DNA molecules for preparing the inventive polypeptides.</p>		

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COMPOUNDS AND METHODS FOR THERAPY AND DIAGNOSIS OF LUNG CANCER

TECHNICAL FIELD

5 The present invention relates generally to therapy and diagnosis of cancer, such as lung cancer. The invention is more specifically related to polypeptides comprising at least a portion of a lung tumor protein, and to polynucleotides encoding such polypeptides. Such polypeptides and polynucleotides may be used in vaccines and pharmaceutical compositions for prevention and treatment of lung cancer, and for the
10 diagnosis and monitoring of such cancers.

BACKGROUND OF THE INVENTION

Lung cancer is the primary cause of cancer death among both men and women in the U.S., with an estimated 172,000 new cases being reported in 1994. The five-year survival rate among all lung cancer patients, regardless of the stage of disease
15 at diagnosis, is only 13%. This contrasts with a five-year survival rate of 46% among cases detected while the disease is still localized. However, only 16% of lung cancers are discovered before the disease has spread.

Early detection is difficult since clinical symptoms are often not seen until the disease has reached an advanced stage. Currently, diagnosis is aided by the
20 use of chest x-rays, analysis of the type of cells contained in sputum and fiberoptic examination of the bronchial passages. Treatment regimens are determined by the type and stage of the cancer, and include surgery, radiation therapy and/or chemotherapy. In spite of considerable research into therapies for the disease, lung cancer remains difficult to treat.

25 Accordingly, there remains a need in the art for improved vaccines, treatment methods and diagnostic techniques for lung cancer.

SUMMARY OF THE INVENTION

Briefly stated, the present invention provides compositions and methods for the diagnosis and therapy of cancer, such as lung cancer. In one aspect, the present

invention provides polypeptides comprising at least a portion of a lung tumor protein, or a variant thereof. Certain portions and other variants are immunogenic, such that the ability of the variant to react with antigen-specific antisera is not substantially diminished. Within certain embodiments, the polypeptide comprises a sequence that is encoded by a polynucleotide sequence selected from the group consisting of: (a) sequences recited in any one of SEQ ID NO: 1-3, 6-8, 10-13, 15-27, 29, 30, 32, 34-49, 51, 52, 54, 55, 57-59, 61-69, 71, 73, 74, 77, 78, 80-82, 84, 86-96, 107-109, 111, 113, 125, 127, 128, 129, 131-133, 142, 144, 148-151, 153, 154, 157, 158, 160, 167, 168, 171, 179, 182, 184-186, 188-191, 193, 194, 198-207, 209, 210, 213, 214, 217, 220-224, 253-337, 345, 347 and 349; (b) variants of a sequence recited in any one of SEQ ID NO: 1-3, 6-8, 10-13, 15-27, 29, 30, 32, 34-49, 51, 52, 54, 55, 57-59, 61-69, 71, 73, 74, 77, 78, 80-82, 84, 86-96, 107-109, 111, 113, 125, 127, 128, 129, 131-133, 142, 144, 148-151, 153, 154, 157, 158, 160, 167, 168, 171, 179, 182, 184-186, 188-191, 193, 194, 198-207, 209, 210, 213, 214, 217, 220-224, 253-337, 345, 347 and 349; and (c) complements of a sequence of (a) or (b). In specific embodiments, the polypeptides of the present invention comprise at least a portion of a tumor protein that includes an amino acid sequence selected from the group consisting of sequences recited in any one of SEQ ID NO: 152, 155, 156, 165, 166, 169, 170, 172, 174, 176, 226-252, 338-344 and 346, and variants thereof.

The present invention further provides polynucleotides that encode a polypeptide as described above, or a portion thereof (such as a portion encoding at least 15 amino acid residues of a lung tumor protein), expression vectors comprising such polynucleotides and host cells transformed or transfected with such expression vectors.

Within other aspects, the present invention provides pharmaceutical compositions comprising a polypeptide or polynucleotide as described above and a physiologically acceptable carrier.

Within a related aspect of the present invention, vaccines for prophylactic or therapeutic use are provided. Such vaccines comprise a polypeptide or polynucleotide as described above and an immunostimulant.

The present invention further provides pharmaceutical compositions that comprise: (a) an antibody or antigen-binding fragment thereof that specifically binds to a lung tumor protein; and (b) a physiologically acceptable carrier.

Within further aspects, the present invention provides pharmaceutical compositions comprising: (a) an antigen presenting cell that expresses a polypeptide as described above and (b) a pharmaceutically acceptable carrier or excipient. Antigen presenting cells include dendritic cells, macrophages, monocytes, fibroblasts and B cells.

Within related aspects, vaccines are provided that comprise: (a) an antigen presenting cell that expresses a polypeptide as described above, and (b) an immunostimulant.

The present invention further provides, in other aspects, fusion proteins that comprise at least one polypeptide as described above, as well as polynucleotides encoding such fusion proteins.

Within related aspects, pharmaceutical compositions comprising a fusion protein, or a polynucleotide encoding a fusion protein, in combination with a physiologically acceptable carrier are provided.

Vaccines are further provided, within other aspects, that comprise a fusion protein, or a polynucleotide encoding a fusion protein, in combination with an immunostimulant.

Within further aspects, the present invention provides methods for inhibiting the development of a cancer in a patient, comprising administering to a patient a pharmaceutical composition or vaccine as recited above.

The present invention further provides, within other aspects, methods for removing tumor cells from a biological sample, comprising contacting a biological sample with T cells that specifically react with a lung tumor protein, wherein the step of contacting is performed under conditions and for a time sufficient to permit the removal of cells expressing the protein from the sample.

Within related aspects, methods are provided for inhibiting the development of a cancer in a patient, comprising administering to a patient a biological sample treated as described above.

Methods are further provided, within other aspects, for stimulating and/or expanding T cells specific for a lung tumor protein, comprising contacting T cells with one or more of: (i) a polypeptide as described above; (ii) a polynucleotide encoding such a polypeptide; and/or (iii) an antigen presenting cell that expresses such a polypeptide; under conditions and for a time sufficient to permit the stimulation and/or expansion of T cells. Determined T cell populations comprising T cells prepared as described above are also provided.

Within further aspects, the present invention provides methods for inhibiting the development of a cancer in a patient, comprising administering to a patient an effective amount of a T cell population as described above.

The present invention further provides methods for inhibiting the development of a cancer in a patient, comprising the steps of: (a) incubating CD4⁺ and/or CD8⁺ T cells determined from a patient with one or more of: (i) a polypeptide comprising at least an immunogenic portion of a lung tumor protein; (ii) a polynucleotide encoding such a polypeptide; and (iii) an antigen-presenting cell that expressed such a polypeptide; and (b) administering to the patient an effective amount of the proliferated T cells, and thereby inhibiting the development of a cancer in the patient. Proliferated cells may, but need not, be cloned prior to administration to the patient.

Within further aspects, the present invention provides methods for determining the presence or absence of a cancer in a patient, comprising: (a) contacting a biological sample obtained from a patient with a binding agent that binds to a polypeptide as recited above; (b) detecting in the sample an amount of polypeptide that binds to the binding agent; and (c) comparing the amount of polypeptide with a predetermined cut-off value, and therefrom determining the presence or absence of a cancer in the patient. Within preferred embodiments, the binding agent is an antibody, more preferably a monoclonal antibody. The cancer may be lung cancer.

The present invention also provides, within other aspects, methods for monitoring the progression of a cancer in a patient. Such methods comprise the steps of: (a) contacting a biological sample obtained from a patient at a first point in time with a binding agent that binds to a polypeptide as recited above; (b) detecting in the

sample an amount of polypeptide that binds to the binding agent; (c) repeating steps (a) and (b) using a biological sample obtained from the patient at a subsequent point in time; and (d) comparing the amount of polypeptide detected in step (c) with the amount detected in step (b) and therefrom monitoring the progression of the cancer in the patient.

The present invention further provides, within other aspects, methods for determining the presence or absence of a cancer in a patient, comprising the steps of: (a) contacting a biological sample obtained from a patient with an oligonucleotide that hybridizes to a polynucleotide that encodes a lung tumor protein; (b) detecting in the sample a level of a polynucleotide, preferably mRNA, that hybridizes to the oligonucleotide; and (c) comparing the level of polynucleotide that hybridizes to the oligonucleotide with a predetermined cut-off value, and therefrom determining the presence or absence of a cancer in the patient. Within certain embodiments, the amount of mRNA is detected via polymerase chain reaction using, for example, at least one oligonucleotide primer that hybridizes to a polynucleotide encoding a polypeptide as recited above, or a complement of such a polynucleotide. Within other embodiments, the amount of mRNA is detected using a hybridization technique, employing an oligonucleotide probe that hybridizes to a polynucleotide that encodes a polypeptide as recited above, or a complement of such a polynucleotide.

In related aspects, methods are provided for monitoring the progression of a cancer in a patient, comprising the steps of: (a) contacting a biological sample obtained from a patient with an oligonucleotide that hybridizes to a polynucleotide that encodes a lung tumor protein; (b) detecting in the sample an amount of a polynucleotide that hybridizes to the oligonucleotide; (c) repeating steps (a) and (b) using a biological sample obtained from the patient at a subsequent point in time; and (d) comparing the amount of polynucleotide detected in step (c) with the amount detected in step (b) and therefrom monitoring the progression of the cancer in the patient.

Within further aspects, the present invention provides antibodies, such as monoclonal antibodies, that bind to a polypeptide as described above, as well as diagnostic kits comprising such antibodies. Diagnostic kits comprising one or more oligonucleotide probes or primers as described above are also provided.

These and other aspects of the present invention will become apparent upon reference to the following detailed description and attached drawings. All references disclosed herein are hereby incorporated by reference in their entirety as if each was incorporated individually.

5

SEQUENCE IDENTIFIERS

- SEQ ID NO: 1 is the determined cDNA sequence for LST-S1-2
- SEQ ID NO: 2 is the determined cDNA sequence for LST-S1-28
- SEQ ID NO: 3 is the determined cDNA sequence for LST-S1-90
- 10 SEQ ID NO: 4 is the determined cDNA sequence for LST-S1-144
- SEQ ID NO: 5 is the determined cDNA sequence for LST-S1-133
- SEQ ID NO: 6 is the determined cDNA sequence for LST-S1-169
- SEQ ID NO: 7 is the determined cDNA sequence for LST-S2-6
- SEQ ID NO: 8 is the determined cDNA sequence for LST-S2-11
- 15 SEQ ID NO: 9 is the determined cDNA sequence for LST-S2-17
- SEQ ID NO: 10 is the determined cDNA sequence for LST-S2-25
- SEQ ID NO: 11 is the determined cDNA sequence for LST-S2-39
- SEQ ID NO: 12 is a first determined cDNA sequence for LST-S2-43
- SEQ ID NO: 13 is a second determined cDNA sequence for LST-S2-43
- 20 SEQ ID NO: 14 is the determined cDNA sequence for LST-S2-65
- SEQ ID NO: 15 is the determined cDNA sequence for LST-S2-68
- SEQ ID NO: 16 is the determined cDNA sequence for LST-S2-72
- SEQ ID NO: 17 is the determined cDNA sequence for LST-S2-74
- SEQ ID NO: 18 is the determined cDNA sequence for LST-S2-103
- 25 SEQ ID NO: 19 is the determined cDNA sequence for LST-S2-N1-1F
- SEQ ID NO: 20 is the determined cDNA sequence for LST-S2-N1-2A
- SEQ ID NO: 21 is the determined cDNA sequence for LST-S2-N1-4H
- SEQ ID NO: 22 is the determined cDNA sequence for LST-S2-N1-5A
- SEQ ID NO: 23 is the determined cDNA sequence for LST-S2-N1-6B
- 30 SEQ ID NO: 24 is the determined cDNA sequence for LST-S2-N1-7B
- SEQ ID NO: 25 is the determined cDNA sequence for LST-S2-N1-7H

- SEQ ID NO: 26 is the determined cDNA sequence for LST-S2-N1-8A
- SEQ ID NO: 27 is the determined cDNA sequence for LST-S2-N1-8D
- SEQ ID NO: 28 is the determined cDNA sequence for LST-S2-N1-9A
- SEQ ID NO: 29 is the determined cDNA sequence for LST-S2-N1-9E
- 5 SEQ ID NO: 30 is the determined cDNA sequence for LST-S2-N1-10A
- SEQ ID NO: 31 is the determined cDNA sequence for LST-S2-N1-10G
- SEQ ID NO: 32 is the determined cDNA sequence for LST-S2-N1-11A
- SEQ ID NO: 33 is the determined cDNA sequence for LST-S2-N1-12C
- SEQ ID NO: 34 is the determined cDNA sequence for LST-S2-N1-12E
- 10 SEQ ID NO: 35 is the determined cDNA sequence for LST-S2-B1-3D
- SEQ ID NO: 36 is the determined cDNA sequence for LST-S2-B1-6C
- SEQ ID NO: 37 is the determined cDNA sequence for LST-S2-B1-5D
- SEQ ID NO: 38 is the determined cDNA sequence for LST-S2-B1-5F
- SEQ ID NO: 39 is the determined cDNA sequence for LST-S2-B1-6G
- 15 SEQ ID NO: 40 is the determined cDNA sequence for LST-S2-B1-8A
- SEQ ID NO: 41 is the determined cDNA sequence for LST-S2-B1-8D
- SEQ ID NO: 42 is the determined cDNA sequence for LST-S2-B1-10A
- SEQ ID NO: 43 is the determined cDNA sequence for LST-S2-B1-9B
- SEQ ID NO: 44 is the determined cDNA sequence for LST-S2-B1-9F
- 20 SEQ ID NO: 45 is the determined cDNA sequence for LST-S2-B1-12D
- SEQ ID NO: 46 is the determined cDNA sequence for LST-S2-I2-2B
- SEQ ID NO: 47 is the determined cDNA sequence for LST-S2-I2-5F
- SEQ ID NO: 48 is the determined cDNA sequence for LST-S2-I2-6B
- SEQ ID NO: 49 is the determined cDNA sequence for LST-S2-I2-7F
- 25 SEQ ID NO: 50 is the determined cDNA sequence for LST-S2-I2-8G
- SEQ ID NO: 51 is the determined cDNA sequence for LST-S2-I2-9E
- SEQ ID NO: 52 is the determined cDNA sequence for LST-S2-I2-12B
- SEQ ID NO: 53 is the determined cDNA sequence for LST-S2-H2-2C
- SEQ ID NO: 54 is the determined cDNA sequence for LST-S2-H2-1G
- 30 SEQ ID NO: 55 is the determined cDNA sequence for LST-S2-H2-4G
- SEQ ID NO: 56 is the determined cDNA sequence for LST-S2-H2-3H

- SEQ ID NO: 57 is the determined cDNA sequence for LST-S2-H2-5G
- SEQ ID NO: 58 is the determined cDNA sequence for LST-S2-H2-9B
- SEQ ID NO: 59 is the determined cDNA sequence for LST-S2-H2-10H
- SEQ ID NO: 60 is the determined cDNA sequence for LST-S2-H2-12D
- 5 SEQ ID NO: 61 is the determined cDNA sequence for LST-S3-2
- SEQ ID NO: 62 is the determined cDNA sequence for LST-S3-4
- SEQ ID NO: 63 is the determined cDNA sequence for LST-S3-7
- SEQ ID NO: 64 is the determined cDNA sequence for LST-S3-8
- SEQ ID NO: 65 is the determined cDNA sequence for LST-S3-12
- 10 SEQ ID NO: 66 is the determined cDNA sequence for LST-S3-13
- SEQ ID NO: 67 is the determined cDNA sequence for LST-S3-14
- SEQ ID NO: 68 is the determined cDNA sequence for LST-S3-16
- SEQ ID NO: 69 is the determined cDNA sequence for LST-S3-21
- SEQ ID NO: 70 is the determined cDNA sequence for LST-S3-22
- 15 SEQ ID NO: 71 is the determined cDNA sequence for LST-S1-7
- SEQ ID NO: 72 is the determined cDNA sequence for LST-S1-A-1E
- SEQ ID NO: 73 is the determined cDNA sequence for LST-S1-A-1G
- SEQ ID NO: 74 is the determined cDNA sequence for LST-S1-A-3E
- SEQ ID NO: 75 is the determined cDNA sequence for LST-S1-A-4E
- 20 SEQ ID NO: 76 is the determined cDNA sequence for LST-S1-A-6D
- SEQ ID NO: 77 is the determined cDNA sequence for LST-S1-A-8D
- SEQ ID NO: 78 is the determined cDNA sequence for LST-S1-A-10A
- SEQ ID NO: 79 is the determined cDNA sequence for LST-S1-A-10C
- SEQ ID NO: 80 is the determined cDNA sequence for LST-S1-A-9D
- 25 SEQ ID NO: 81 is the determined cDNA sequence for LST-S1-A-10D
- SEQ ID NO: 82 is the determined cDNA sequence for LST-S1-A-9H
- SEQ ID NO: 83 is the determined cDNA sequence for LST-S1-A-11D
- SEQ ID NO: 84 is the determined cDNA sequence for LST-S1-A-12D
- SEQ ID NO: 85 is the determined cDNA sequence for LST-S1-A-11E
- 30 SEQ ID NO: 86 is the determined cDNA sequence for LST-S1-A-12E
- SEQ ID NO: 87 is the determined cDNA sequence for L513S (T3).

- SEQ ID NO: 88 is the determined cDNA sequence for L513S contig 1.
- SEQ ID NO: 89 is a first determined cDNA sequence for L514S.
- SEQ ID NO: 90 is a second determined cDNA sequence for L514S.
- SEQ ID NO: 91 is a first determined cDNA sequence for L516S.
- 5 SEQ ID NO: 92 is a second determined cDNA sequence for L516S.
- SEQ ID NO: 93 is the determined cDNA sequence for L517S.
- SEQ ID NO: 94 is the extended cDNA sequence for LST-S1-169 (also known as L519S).
- SEQ ID NO: 95 is a first determined cDNA sequence for L520S.
- 10 SEQ ID NO: 96 is a second determined cDNA sequence for L520S.
- SEQ ID NO: 97 is a first determined cDNA sequence for L521S.
- SEQ ID NO: 98 is a second determined cDNA sequence for L521S.
- SEQ ID NO: 99 is the determined cDNA sequence for L522S.
- SEQ ID NO: 100 is the determined cDNA sequence for L523S.
- 15 SEQ ID NO: 101 is the determined cDNA sequence for L524S.
- SEQ ID NO: 102 is the determined cDNA sequence for L525S.
- SEQ ID NO: 103 is the determined cDNA sequence for L526S.
- SEQ ID NO: 104 is the determined cDNA sequence for L527S.
- SEQ ID NO: 105 is the determined cDNA sequence for L528S.
- 20 SEQ ID NO: 106 is the determined cDNA sequence for L529S.
- SEQ ID NO: 107 is a first determined cDNA sequence for L530S.
- SEQ ID NO: 108 is a second determined cDNA sequence for L530S.
- SEQ ID NO: 109 is the determined full-length cDNA sequence for L531S short form.
- SEQ ID NO: 110 is the predicted amino acid sequence encoded by SEQ ID NO: 109.
- 25 SEQ ID NO: 111 is the determined full-length cDNA sequence for L531S long form.
- SEQ ID NO: 112 is the predicted amino acid sequence encoded by SEQ ID NO: 111.
- SEQ ID NO: 113 is the determined full-length cDNA sequence for L520S.
- SEQ ID NO: 114 is the predicted amino acid sequence encoded by SEQ ID NO: 113.
- SEQ ID NO: 115 is the determined cDNA sequence for contig 1.
- 30 SEQ ID NO: 116 is the determined cDNA sequence for contig 3.
- SEQ ID NO: 117 is the determined cDNA sequence for contig 4.

- SEQ ID NO: 118 is the determined cDNA sequence for contig 5.
- SEQ ID NO: 119 is the determined cDNA sequence for contig 7.
- SEQ ID NO: 120 is the determined cDNA sequence for contig 8.
- SEQ ID NO: 121 is the determined cDNA sequence for contig 9.
- 5 SEQ ID NO: 122 is the determined cDNA sequence for contig 10.
- SEQ ID NO: 123 is the determined cDNA sequence for contig 12.
- SEQ ID NO: 124 is the determined cDNA sequence for contig 11.
- SEQ ID NO: 125 is the determined cDNA sequence for contig 13.
- SEQ ID NO: 126 is the determined cDNA sequence for contig 15.
- 10 SEQ ID NO: 127 is the determined cDNA sequence for contig 16.
- SEQ ID NO: 128 is the determined cDNA sequence for contig 17.
- SEQ ID NO: 129 is the determined cDNA sequence for contig 19.
- SEQ ID NO: 130 is the determined cDNA sequence for contig 20.
- SEQ ID NO: 131 is the determined cDNA sequence for contig 22.
- 15 SEQ ID NO: 132 is the determined cDNA sequence for contig 24.
- SEQ ID NO: 133 is the determined cDNA sequence for contig 29.
- SEQ ID NO: 134 is the determined cDNA sequence for contig 31.
- SEQ ID NO: 135 is the determined cDNA sequence for contig 33.
- SEQ ID NO: 136 is the determined cDNA sequence for contig 38.
- 20 SEQ ID NO: 137 is the determined cDNA sequence for contig 39.
- SEQ ID NO: 138 is the determined cDNA sequence for contig 41.
- SEQ ID NO: 139 is the determined cDNA sequence for contig 43.
- SEQ ID NO: 140 is the determined cDNA sequence for contig 44.
- SEQ ID NO: 141 is the determined cDNA sequence for contig 45.
- 25 SEQ ID NO: 142 is the determined cDNA sequence for contig 47.
- SEQ ID NO: 143 is the determined cDNA sequence for contig 48.
- SEQ ID NO: 144 is the determined cDNA sequence for contig 49.
- SEQ ID NO: 145 is the determined cDNA sequence for contig 50.
- SEQ ID NO: 146 is the determined cDNA sequence for contig 53.
- 30 SEQ ID NO: 147 is the determined cDNA sequence for contig 54.
- SEQ ID NO: 148 is the determined cDNA sequence for contig 56.

- SEQ ID NO: 149 is the determined cDNA sequence for contig 57.
- SEQ ID NO: 150 is the determined cDNA sequence for contig 58.
- SEQ ID NO: 151 is the full-length cDNA sequence for L530S.
- SEQ ID NO: 152 is the amino acid sequence encoded by SEQ ID NO: 151.
- 5 SEQ ID NO: 153 is the full-length cDNA sequence of a first variant of L514S.
- SEQ ID NO: 154 is the full-length cDNA sequence of a second variant of L514S.
- SEQ ID NO: 155 is the amino acid sequence encoded by SEQ ID NO: 153.
- SEQ ID NO: 156 is the amino acid sequence encoded by SEQ ID NO: 154.
- SEQ ID NO: 157 is the determined cDNA sequence for contig 59.
- 10 SEQ ID NO: 158 is the full-length cDNA sequence for L763P (also referred to as contig 22).
- SEQ ID NO: 159 is the amino acid sequence encoded by SEQ ID NO: 158.
- SEQ ID NO: 160 is the full-length cDNA sequence for L762P (also referred to as contig 17).
- 15 SEQ ID NO: 161 is the amino acid sequence encoded by SEQ ID NO: 160.
- SEQ ID NO: 162 is the determined cDNA sequence for L515S.
- SEQ ID NO: 163 is the full-length cDNA sequence of a first variant of L524S.
- SEQ ID NO: 164 is the full-length cDNA sequence of a second variant of L524S.
- SEQ ID NO: 165 is the amino acid sequence encoded by SEQ ID NO: 163.
- 20 SEQ ID NO: 166 is the amino acid sequence encoded by SEQ ID NO: 164.
- SEQ ID NO: 167 is the full-length cDNA sequence of a first variant of L762P.
- SEQ ID NO: 168 is the full-length cDNA sequence of a second variant of L762P.
- SEQ ID NO: 169 is the amino acid sequence encoded by SEQ ID NO: 167.
- SEQ ID NO: 170 is the amino acid sequence encoded by SEQ ID NO: 168.
- 25 SEQ ID NO: 171 is the full-length cDNA sequence for L773P (also referred to as contig 56).
- SEQ ID NO: 172 is the amino acid sequence encoded by SEQ ID NO: 171.
- SEQ ID NO: 173 is an extended cDNA sequence for L519S.
- SEQ ID NO: 174 is the predicted amino acid sequence encoded by SEQ ID NO: 174.
- 30 SEQ ID NO: 175 is the full-length cDNA sequence for L523S.
- SEQ ID NO: 176 is the predicted amino acid sequence encoded by SEQ ID NO: 175.

- SEQ ID NO: 177 is the determined cDNA sequence for LST-sub5-7A.
- SEQ ID NO: 178 is the determined cDNA sequence for LST-sub5-8G.
- SEQ ID NO: 179 is the determined cDNA sequence for LST-sub5-8H.
- SEQ ID NO: 180 is the determined cDNA sequence for LST-sub5-10B.
- 5 SEQ ID NO: 181 is the determined cDNA sequence for LST-sub5-10H.
- SEQ ID NO: 182 is the determined cDNA sequence for LST-sub5-12B.
- SEQ ID NO: 183 is the determined cDNA sequence for LST-sub5-11C.
- SEQ ID NO: 184 is the determined cDNA sequence for LST-sub6-1c.
- SEQ ID NO: 185 is the determined cDNA sequence for LST-sub6-2f.
- 10 SEQ ID NO: 186 is the determined cDNA sequence for LST-sub6-2G.
- SEQ ID NO: 187 is the determined cDNA sequence for LST-sub6-4d.
- SEQ ID NO: 188 is the determined cDNA sequence for LST-sub6-4e.
- SEQ ID NO: 189 is the determined cDNA sequence for LST-sub6-4f.
- SEQ ID NO: 190 is the determined cDNA sequence for LST-sub6-3h.
- 15 SEQ ID NO: 191 is the determined cDNA sequence for LST-sub6-5d.
- SEQ ID NO: 192 is the determined cDNA sequence for LST-sub6-5h.
- SEQ ID NO: 193 is the determined cDNA sequence for LST-sub6-6h.
- SEQ ID NO: 194 is the determined cDNA sequence for LST-sub6-7a.
- SEQ ID NO: 195 is the determined cDNA sequence for LST-sub6-8a.
- 20 SEQ ID NO: 196 is the determined cDNA sequence for LST-sub6-7d.
- SEQ ID NO: 197 is the determined cDNA sequence for LST-sub6-7e.
- SEQ ID NO: 198 is the determined cDNA sequence for LST-sub6-8e.
- SEQ ID NO: 199 is the determined cDNA sequence for LST-sub6-7g.
- SEQ ID NO: 200 is the determined cDNA sequence for LST-sub6-9f.
- 25 SEQ ID NO: 201 is the determined cDNA sequence for LST-sub6-9h.
- SEQ ID NO: 202 is the determined cDNA sequence for LST-sub6-11b.
- SEQ ID NO: 203 is the determined cDNA sequence for LST-sub6-11c.
- SEQ ID NO: 204 is the determined cDNA sequence for LST-sub6-12c.
- SEQ ID NO: 205 is the determined cDNA sequence for LST-sub6-12e.
- 30 SEQ ID NO: 206 is the determined cDNA sequence for LST-sub6-12f.
- SEQ ID NO: 207 is the determined cDNA sequence for LST-sub6-11g.

- SEQ ID NO: 208 is the determined cDNA sequence for LST-sub6-12g.
- SEQ ID NO: 209 is the determined cDNA sequence for LST-sub6-12h.
- SEQ ID NO: 210 is the determined cDNA sequence for LST-sub6-II-1a.
- SEQ ID NO: 211 is the determined cDNA sequence for LST-sub6-II-2b.
- 5 SEQ ID NO: 212 is the determined cDNA sequence for LST-sub6-II-2g.
- SEQ ID NO: 213 is the determined cDNA sequence for LST-sub6-II-1h.
- SEQ ID NO: 214 is the determined cDNA sequence for LST-sub6-II-4a.
- SEQ ID NO: 215 is the determined cDNA sequence for LST-sub6-II-4b.
- SEQ ID NO: 216 is the determined cDNA sequence for LST-sub6-II-3e.
- 10 SEQ ID NO: 217 is the determined cDNA sequence for LST-sub6-II-4f.
- SEQ ID NO: 218 is the determined cDNA sequence for LST-sub6-II-4g.
- SEQ ID NO: 219 is the determined cDNA sequence for LST-sub6-II-4h.
- SEQ ID NO: 220 is the determined cDNA sequence for LST-sub6-II-5c.
- SEQ ID NO: 221 is the determined cDNA sequence for LST-sub6-II-5e.
- 15 SEQ ID NO: 222 is the determined cDNA sequence for LST-sub6-II-6f.
- SEQ ID NO: 223 is the determined cDNA sequence for LST-sub6-II-5g.
- SEQ ID NO: 224 is the determined cDNA sequence for LST-sub6-II-6g.
- SEQ ID NO: 225 is the amino acid sequence for L528S.
- SEQ ID NO: 226-251 are synthetic peptides derived from L762P.
- 20 SEQ ID NO: 252 is the expressed amino acid sequence of L514S.
- SEQ ID NO: 253 is the DNA sequence corresponding to SEQ ID NO: 252.
- SEQ ID NO: 254 is the DNA sequence of a L762P expression construct.
- SEQ ID NO: 255 is the determined cDNA sequence for clone 23785.
- SEQ ID NO: 256 is the determined cDNA sequence for clone 23786.
- 25 SEQ ID NO: 257 is the determined cDNA sequence for clone 23788.
- SEQ ID NO: 258 is the determined cDNA sequence for clone 23790.
- SEQ ID NO: 259 is the determined cDNA sequence for clone 23793.
- SEQ ID NO: 260 is the determined cDNA sequence for clone 23794.
- SEQ ID NO: 261 is the determined cDNA sequence for clone 23795.
- 30 SEQ ID NO: 262 is the determined cDNA sequence for clone 23796.
- SEQ ID NO: 263 is the determined cDNA sequence for clone 23797.

- SEQ ID NO: 264 is the determined cDNA sequence for clone 23798.
- SEQ ID NO: 265 is the determined cDNA sequence for clone 23799.
- SEQ ID NO: 266 is the determined cDNA sequence for clone 23800.
- SEQ ID NO: 267 is the determined cDNA sequence for clone 23802.
- 5 SEQ ID NO: 268 is the determined cDNA sequence for clone 23803.
- SEQ ID NO: 269 is the determined cDNA sequence for clone 23804.
- SEQ ID NO: 270 is the determined cDNA sequence for clone 23805.
- SEQ ID NO: 271 is the determined cDNA sequence for clone 23806.
- SEQ ID NO: 272 is the determined cDNA sequence for clone 23807.
- 10 SEQ ID NO: 273 is the determined cDNA sequence for clone 23808.
- SEQ ID NO: 274 is the determined cDNA sequence for clone 23809.
- SEQ ID NO: 275 is the determined cDNA sequence for clone 23810.
- SEQ ID NO: 276 is the determined cDNA sequence for clone 23811.
- SEQ ID NO: 277 is the determined cDNA sequence for clone 23812.
- 15 SEQ ID NO: 278 is the determined cDNA sequence for clone 23813.
- SEQ ID NO: 279 is the determined cDNA sequence for clone 23815.
- SEQ ID NO: 280 is the determined cDNA sequence for clone 25298.
- SEQ ID NO: 281 is the determined cDNA sequence for clone 25299.
- SEQ ID NO: 282 is the determined cDNA sequence for clone 25300.
- 20 SEQ ID NO: 283 is the determined cDNA sequence for clone 25301.
- SEQ ID NO: 284 is the determined cDNA sequence for clone 25304.
- SEQ ID NO: 285 is the determined cDNA sequence for clone 25309.
- SEQ ID NO: 286 is the determined cDNA sequence for clone 25312.
- SEQ ID NO: 287 is the determined cDNA sequence for clone 25317.
- 25 SEQ ID NO: 288 is the determined cDNA sequence for clone 25321.
- SEQ ID NO: 289 is the determined cDNA sequence for clone 25323.
- SEQ ID NO: 290 is the determined cDNA sequence for clone 25327.
- SEQ ID NO: 291 is the determined cDNA sequence for clone 25328.
- SEQ ID NO: 292 is the determined cDNA sequence for clone 25332.
- 30 SEQ ID NO: 293 is the determined cDNA sequence for clone 25333.
- SEQ ID NO: 294 is the determined cDNA sequence for clone 25336.

- SEQ ID NO: 295 is the determined cDNA sequence for clone 25340.
- SEQ ID NO: 296 is the determined cDNA sequence for clone 25342.
- SEQ ID NO: 297 is the determined cDNA sequence for clone 25356.
- SEQ ID NO: 298 is the determined cDNA sequence for clone 25357.
- 5 SEQ ID NO: 299 is the determined cDNA sequence for clone 25361.
- SEQ ID NO: 300 is the determined cDNA sequence for clone 25363.
- SEQ ID NO: 301 is the determined cDNA sequence for clone 25397.
- SEQ ID NO: 302 is the determined cDNA sequence for clone 25402.
- SEQ ID NO: 303 is the determined cDNA sequence for clone 25403.
- 10 SEQ ID NO: 304 is the determined cDNA sequence for clone 25405.
- SEQ ID NO: 305 is the determined cDNA sequence for clone 25407.
- SEQ ID NO: 306 is the determined cDNA sequence for clone 25409.
- SEQ ID NO: 307 is the determined cDNA sequence for clone 25396.
- SEQ ID NO: 308 is the determined cDNA sequence for clone 25414.
- 15 SEQ ID NO: 309 is the determined cDNA sequence for clone 25410.
- SEQ ID NO: 310 is the determined cDNA sequence for clone 25406.
- SEQ ID NO: 311 is the determined cDNA sequence for clone 25306.
- SEQ ID NO: 312 is the determined cDNA sequence for clone 25362.
- SEQ ID NO: 313 is the determined cDNA sequence for clone 25360.
- 20 SEQ ID NO: 314 is the determined cDNA sequence for clone 25398.
- SEQ ID NO: 315 is the determined cDNA sequence for clone 25355.
- SEQ ID NO: 316 is the determined cDNA sequence for clone 25351.
- SEQ ID NO: 317 is the determined cDNA sequence for clone 25331.
- SEQ ID NO: 318 is the determined cDNA sequence for clone 25338.
- 25 SEQ ID NO: 319 is the determined cDNA sequence for clone 25335.
- SEQ ID NO: 320 is the determined cDNA sequence for clone 25329.
- SEQ ID NO: 321 is the determined cDNA sequence for clone 25324.
- SEQ ID NO: 322 is the determined cDNA sequence for clone 25322.
- SEQ ID NO: 323 is the determined cDNA sequence for clone 25319.
- 30 SEQ ID NO: 324 is the determined cDNA sequence for clone 25316.
- SEQ ID NO: 325 is the determined cDNA sequence for clone 25311.

- SEQ ID NO: 326 is the determined cDNA sequence for clone 25310.
- SEQ ID NO: 327 is the determined cDNA sequence for clone 25302.
- SEQ ID NO: 328 is the determined cDNA sequence for clone 25315.
- SEQ ID NO: 329 is the determined cDNA sequence for clone 25308.
- 5 SEQ ID NO: 330 is the determined cDNA sequence for clone 25303.
- SEQ ID NO: 331-337 are the cDNA sequences of isoforms of the p53 tumor suppressor homologue, p63 (also referred to as L530S).
- SEQ ID NO: 338-344 are the amino acid sequences encoded by SEQ ID NO: 331-337, respectively.
- 10 SEQ ID NO: 345 is a second cDNA sequence for the antigen L763P.
- SEQ ID NO: 346 is the amino acid sequence encoded by the sequence of SEQ ID NO: 345.
- SEQ ID NO: 347 is a determined full-length cDNA sequence for L523S.
- SEQ ID NO: 348 is the predicted amino acid sequence encoded by SEQ ID NO: 347.
- 15 SEQ ID NO: 349 is the cDNA sequence encoding the N-terminal portion of L773P.
- SEQ ID NO: 350 is the amino acid sequence of the N-terminal portion of L773P.

DETAILED DESCRIPTION OF THE INVENTION

- As noted above, the present invention is generally directed to
- 20 compositions and methods for the therapy and diagnosis of cancer, such as lung cancer. The compositions described herein may include lung tumor polypeptides, polynucleotides encoding such polypeptides, binding agents such as antibodies, antigen presenting cells (APCs) and/or immune system cells (e.g., T cells). Polypeptides of the present invention generally comprise at least a portion (such as an immunogenic
- 25 portion) of a lung tumor protein or a variant thereof. A "lung tumor protein" is a protein that is expressed in lung tumor cells at a level that is at least two fold, and preferably at least five fold, greater than the level of expression in a normal tissue, as determined using a representative assay provided herein. Certain lung tumor proteins are tumor proteins that react detectably (within an immunoassay, such as an ELISA or Western
- 30 blot) with antisera of a patient afflicted with lung cancer. Polynucleotides of the subject invention generally comprise a DNA or RNA sequence that encodes all or a portion of

such a polypeptide, or that is complementary to such a sequence. Antibodies are generally immune system proteins, or antigen-binding fragments thereof, that are capable of binding to a polypeptide as described above. Antigen presenting cells include dendritic cells, macrophages, monocytes, fibroblasts and B-cells that express a polypeptide as described above. T cells that may be employed within such compositions are generally T cells that are specific for a polypeptide as described above.

The present invention is based on the discovery human lung tumor proteins. Sequences of polynucleotides encoding specific tumor proteins are provided in SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349.

LUNG TUMOR PROTEIN POLYNUCLEOTIDES

Any polynucleotide that encodes a lung tumor protein or a portion or other variant thereof as described herein is encompassed by the present invention. Preferred polynucleotides comprise at least 15 consecutive nucleotides, preferably at least 30 consecutive nucleotides and more preferably at least 45 consecutive nucleotides, that encode a portion of a lung tumor protein. More preferably, a polynucleotide encodes an immunogenic portion of a lung tumor protein. Polynucleotides complementary to any such sequences are also encompassed by the present invention. Polynucleotides may be single-stranded (coding or antisense) or double-stranded, and may be DNA (genomic, cDNA or synthetic) or RNA molecules. RNA molecules include HnRNA molecules, which contain introns and correspond to a DNA molecule in a one-to-one manner, and mRNA molecules, which do not contain introns. Additional coding or non-coding sequences may, but need not, be present within a polynucleotide of the present invention, and a polynucleotide may, but need not, be linked to other molecules and/or support materials.

Polynucleotides may comprise a native sequence (*i.e.*, an endogenous sequence that encodes a lung tumor protein or a portion thereof) or may comprise a variant of such a sequence. Polynucleotide variants may contain one or more substitutions, additions, deletions and/or insertions such that the immunogenicity of the

encoded polypeptide is not diminished, relative to a native tumor protein. The effect on the immunogenicity of the encoded polypeptide may generally be assessed as described herein. Variants preferably exhibit at least about 70% identity, more preferably at least about 80% identity and most preferably at least about 90% identity to a polynucleotide sequence that encodes a native lung tumor protein or a portion thereof. The term "variants" also encompasses homologous genes of xenogenic origin.

Two polynucleotide or polypeptide sequences are said to be "identical" if the sequence of nucleotides or amino acids in the two sequences is the same when aligned for maximum correspondence as described below. Comparisons between two sequences are typically performed by comparing the sequences over a comparison window to identify and compare local regions of sequence similarity. A "comparison window" as used herein, refers to a segment of at least about 20 contiguous positions, usually 30 to about 75, 40 to about 50, in which a sequence may be compared to a reference sequence of the same number of contiguous positions after the two sequences are optimally aligned.

Optimal alignment of sequences for comparison may be conducted using the Megalign program in the Lasergene suite of bioinformatics software (DNASTAR, Inc., Madison, WI), using default parameters. This program embodies several alignment schemes described in the following references: Dayhoff, M.O. (1978) A model of evolutionary change in proteins - Matrices for detecting distant relationships. In Dayhoff, M.O. (ed.) Atlas of Protein Sequence and Structure, National Biomedical Research Foundation, Washington DC Vol. 5, Suppl. 3, pp. 345-358; Hein J. (1990) Unified Approach to Alignment and Phylogenies pp. 626-645 *Methods in Enzymology* vol. 183, Academic Press, Inc., San Diego, CA; Higgins, D.G. and Sharp, P.M. (1989) *CABIOS* 5:151-153; Myers, E.W. and Muller W. (1988) *CABIOS* 4:11-17; Robinson, E.D. (1971) *Comb. Theor* 11:105; Santou, N. Nes, M. (1987) *Mol. Biol. Evol.* 4:406-425; Sneath, P.H.A. and Sokal, R.R. (1973) *Numerical Taxonomy - the Principles and Practice of Numerical Taxonomy*, Freeman Press, San Francisco, CA; Wilbur, W.J. and Lipman, D.J. (1983) *Proc. Natl. Acad. Sci. USA* 80:726-730.

Preferably, the "percentage of sequence identity" is determined by comparing two optimally aligned sequences over a window of comparison of at least 20

positions, wherein the portion of the polynucleotide or polypeptide sequence in the comparison window may comprise additions or deletions (i.e. gaps) of 20 percent or less, usually 5 to 15 percent, or 10 to 12 percent, as compared to the reference sequences (which does not comprise additions or deletions) for optimal alignment of the two sequences. The percentage is calculated by determining the number of positions at which the identical nucleic acid bases or amino acid residue occurs in both sequences to yield the number of matched positions, dividing the number of matched positions by the total number of positions in the reference sequence (i.e. the window size) and multiplying the results by 100 to yield the percentage of sequence identity.

10 Variants may also, or alternatively, be substantially homologous to a native gene, or a portion or complement thereof. Such polynucleotide variants are capable of hybridizing under moderately stringent conditions to a naturally occurring DNA sequence encoding a native lung tumor protein (or a complementary sequence). Suitable moderately stringent conditions include prewashing in a solution of 5 X SSC, 0.5% SDS, 1.0 mM EDTA (pH 8.0); hybridizing at 50°C-65°C, 5 X SSC, overnight; 15 followed by washing twice at 65°C for 20 minutes with each of 2X, 0.5X and 0.2X SSC containing 0.1% SDS.

It will be appreciated by those of ordinary skill in the art that, as a result of the degeneracy of the genetic code, there are many nucleotide sequences that encode a polypeptide as described herein. Some of these polynucleotides bear minimal 20 homology to the nucleotide sequence of any native gene. Nonetheless, polynucleotides that vary due to differences in codon usage are specifically contemplated by the present invention. Further, alleles of the genes comprising the polynucleotide sequences provided herein are within the scope of the present invention. Alleles are endogenous 25 genes that are altered as a result of one or more mutations, such as deletions, additions and/or substitutions of nucleotides. The resulting mRNA and protein may, but need not, have an altered structure or function. Alleles may be identified using standard techniques (such as hybridization, amplification and/or database sequence comparison).

Polynucleotides may be prepared using any of a variety of techniques. 30 For example, a polynucleotide may be identified, as described in more detail below, by screening a microarray of cDNAs for tumor-associated expression (i.e., expression that

is at least two fold greater in a lung tumor than in normal tissue, as determined using a representative assay provided herein). Such screens may be performed using a Synteni microarray (Palo Alto, CA) according to the manufacturer's instructions (and essentially as described by Schena et al., *Proc. Natl. Acad. Sci. USA* 93:10614-10619, 1996 and
5 Heller et al., *Proc. Natl. Acad. Sci. USA* 94:2150-2155, 1997). Alternatively, polypeptides may be amplified from cDNA prepared from cells expressing the proteins described herein, such as lung tumor cells. Such polynucleotides may be amplified via polymerase chain reaction (PCR). For this approach, sequence-specific primers may be designed based on the sequences provided herein, and may be purchased or synthesized.

10 An amplified portion may be used to isolate a full length gene from a suitable library (e.g., a lung tumor cDNA library) using well known techniques. Within such techniques, a library (cDNA or genomic) is screened using one or more polynucleotide probes or primers suitable for amplification. Preferably, a library is size-selected to include larger molecules. Random primed libraries may also be
15 preferred for identifying 5' and upstream regions of genes. Genomic libraries are preferred for obtaining introns and extending 5' sequences.

For hybridization techniques, a partial sequence may be labeled (e.g., by nick-translation or end-labeling with ^{32}P) using well known techniques. A bacterial or bacteriophage library is then screened by hybridizing filters containing denatured
20 bacterial colonies (or lawns containing phage plaques) with the labeled probe (see Sambrook et al., *Molecular Cloning: A Laboratory Manual*, Cold Spring Harbor Laboratories, Cold Spring Harbor, NY, 1989). Hybridizing colonies or plaques are selected and expanded, and the DNA is isolated for further analysis. cDNA clones may be analyzed to determine the amount of additional sequence by, for example, PCR using
25 a primer from the partial sequence and a primer from the vector. Restriction maps and partial sequences may be generated to identify one or more overlapping clones. The complete sequence may then be determined using standard techniques, which may involve generating a series of deletion clones. The resulting overlapping sequences are then assembled into a single contiguous sequence. A full length cDNA molecule can be
30 generated by ligating suitable fragments, using well known techniques.

Alternatively, there are numerous amplification techniques for obtaining a full length coding sequence from a partial cDNA sequence. Within such techniques, amplification is generally performed via PCR. Any of a variety of commercially available kits may be used to perform the amplification step. Primers may be designed using, for example, software well known in the art. Primers are preferably 22-30 nucleotides in length, have a GC content of at least 50% and anneal to the target sequence at temperatures of about 68°C to 72°C. The amplified region may be sequenced as described above, and overlapping sequences assembled into a contiguous sequence.

One such amplification technique is inverse PCR (see Triglia et al., *Nucl. Acids Res.* 16:8186, 1988), which uses restriction enzymes to generate a fragment in the known region of the gene. The fragment is then circularized by intramolecular ligation and used as a template for PCR with divergent primers derived from the known region. Within an alternative approach, sequences adjacent to a partial sequence may be retrieved by amplification with a primer to a linker sequence and a primer specific to a known region. The amplified sequences are typically subjected to a second round of amplification with the same linker primer and a second primer specific to the known region. A variation on this procedure, which employs two primers that initiate extension in opposite directions from the known sequence, is described in WO 96/38591. Another such technique is known as "rapid amplification of cDNA ends" or RACE. This technique involves the use of an internal primer and an external primer, which hybridizes to a polyA region or vector sequence, to identify sequences that are 5' and 3' of a known sequence. Additional techniques include capture PCR (Lagerstrom et al., *PCR Methods Applic.* 1:111-19, 1991) and walking PCR (Parker et al., *Nucl. Acids Res.* 19:3055-60, 1991). Other methods employing amplification may also be employed to obtain a full length cDNA sequence.

In certain instances, it is possible to obtain a full length cDNA sequence by analysis of sequences provided in an expressed sequence tag (EST) database, such as that available from GenBank. Searches for overlapping ESTs may generally be performed using well known programs (e.g., NCBI BLAST searches), and such ESTs

may be used to generate a contiguous full length sequence. Full length DNA sequences may also be obtained by analysis of genomic fragments.

Certain nucleic acid sequences of cDNA molecules encoding portions of lung tumor proteins are provided in SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349.

Polynucleotide variants may generally be prepared by any method known in the art, including chemical synthesis by, for example, solid phase phosphoramidite chemical synthesis. Modifications in a polynucleotide sequence may also be introduced using standard mutagenesis techniques, such as oligonucleotide-directed site-specific mutagenesis (see Adelman et al., *DNA* 2:183, 1983). Alternatively, RNA molecules may be generated by *in vitro* or *in vivo* transcription of DNA sequences encoding a lung tumor protein, or portion thereof, provided that the DNA is incorporated into a vector with a suitable RNA polymerase promoter (such as T7 or SP6). Certain portions may be used to prepare an encoded polypeptide, as described herein. In addition, or alternatively, a portion may be administered to a patient such that the encoded polypeptide is generated *in vivo* (e.g., by transfecting antigen-presenting cells, such as dendritic cells, with a cDNA construct encoding a lung tumor polypeptide, and administering the transfected cells to the patient).

A portion of a sequence complementary to a coding sequence (i.e., an antisense polynucleotide) may also be used as a probe or to modulate gene expression. cDNA constructs that can be transcribed into antisense RNA may also be introduced into cells of tissues to facilitate the production of antisense RNA. An antisense polynucleotide may be used, as described herein, to inhibit expression of a tumor protein. Antisense technology can be used to control gene expression through triple-helix formation, which compromises the ability of the double helix to open sufficiently for the binding of polymerases, transcription factors or regulatory molecules (see Gee et al., *In Huber and Carr, Molecular and Immunologic Approaches*, Futura Publishing Co. (Mt. Kisco, NY; 1994)). Alternatively, an antisense molecule may be designed to hybridize with a control region of a gene (e.g., promoter, enhancer or transcription

initiation site), and block transcription of the gene; or to block translation by inhibiting binding of a transcript to ribosomes.

A portion of a coding sequence, or of a complementary sequence, may also be designed as a probe or primer to detect gene expression. Probes may be labeled
5 with a variety of reporter groups, such as radionuclides and enzymes, and are preferably at least 10 nucleotides in length, more preferably at least 20 nucleotides in length and still more preferably at least 30 nucleotides in length. Primers, as noted above, are preferably 22-30 nucleotides in length.

Any polynucleotide may be further modified to increase stability *in vivo*.
10 Possible modifications include, but are not limited to, the addition of flanking sequences at the 5' and/or 3' ends; the use of phosphorothioate or 2' O-methyl rather than phosphodiesterase linkages in the backbone; and/or the inclusion of nontraditional bases such as inosine, queosine and wybutosine, as well as acetyl- methyl-, thio- and other modified forms of adenine, cytidine, guanine, thymine and uridine.

15 Nucleotide sequences as described herein may be joined to a variety of other nucleotide sequences using established recombinant DNA techniques. For example, a polynucleotide may be cloned into any of a variety of cloning vectors, including plasmids, phagemids, lambda phage derivatives and cosmids. Vectors of particular interest include expression vectors, replication vectors, probe generation
20 vectors and sequencing vectors. In general, a vector will contain an origin of replication functional in at least one organism, convenient restriction endonuclease sites and one or more selectable markers. Other elements will depend upon the desired use, and will be apparent to those of ordinary skill in the art.

Within certain embodiments, polynucleotides may be formulated so as to
25 permit entry into a cell of a mammal, and expression therein. Such formulations are particularly useful for therapeutic purposes, as described below. Those of ordinary skill in the art will appreciate that there are many ways to achieve expression of a polynucleotide in a target cell, and any suitable method may be employed. For example, a polynucleotide may be incorporated into a viral vector such as, but not
30 limited to, adenovirus, adeno-associated virus, retrovirus, or vaccinia or other pox virus (e.g., avian pox virus). The polynucleotides may also be administered as naked

plasmid vectors. Techniques for incorporating DNA into such vectors are well known to those of ordinary skill in the art. A retroviral vector may additionally transfer or incorporate a gene for a selectable marker (to aid in the identification or selection of transduced cells) and/or a targeting moiety, such as a gene that encodes a ligand for a receptor on a specific target cell, to render the vector target specific. Targeting may also be accomplished using an antibody, by methods known to those of ordinary skill in the art.

Other formulations for therapeutic purposes include colloidal dispersion systems, such as macromolecule complexes, nanocapsules, microspheres, beads, and lipid-based systems including oil-in-water emulsions, micelles, mixed micelles, and liposomes. A preferred colloidal system for use as a delivery vehicle *in vitro* and *in vivo* is a liposome (*i.e.*, an artificial membrane vesicle). The preparation and use of such systems is well known in the art.

15 LUNG TUMOR POLYPEPTIDES

Within the context of the present invention, polypeptides may comprise at least an immunogenic portion of a lung tumor protein or a variant thereof, as described herein. As noted above, a "lung tumor protein" is a protein that is expressed by lung tumor cells. Proteins that are lung tumor proteins also react detectably within an immunoassay (such as an ELISA) with antisera from a patient with lung cancer. Polypeptides as described herein may be of any length. Additional sequences derived from the native protein and/or heterologous sequences may be present, and such sequences may (but need not) possess further immunogenic or antigenic properties.

An "immunogenic portion," as used herein is a portion of a protein that is recognized (*i.e.*, specifically bound) by a B-cell and/or T-cell surface antigen receptor. Such immunogenic portions generally comprise at least 5 amino acid residues, more preferably at least 10, and still more preferably at least 20 amino acid residues of a lung tumor protein or a variant thereof. Certain preferred immunogenic portions include peptides in which an N-terminal leader sequence and/or transmembrane domain have been deleted. Other preferred immunogenic portions may

contain a small N- and/or C-terminal deletion (e.g., 1-30 amino acids, preferably 5-15 amino acids), relative to the mature protein.

Immunogenic portions may generally be identified using well known techniques, such as those summarized in Paul, *Fundamental Immunology*, 3rd ed., 243-247 (Raven Press, 1993) and references cited therein. Such techniques include screening polypeptides for the ability to react with antigen-specific antibodies, antisera and/or T-cell lines or clones. As used herein, antisera and antibodies are "antigen-specific" if they specifically bind to an antigen (i.e., they react with the protein in an ELISA or other immunoassay, and do not react detectably with unrelated proteins). Such antisera and antibodies may be prepared as described herein, and using well known techniques. An immunogenic portion of a native lung tumor protein is a portion that reacts with such antisera and/or T-cells at a level that is not substantially less than the reactivity of the full length polypeptide (e.g., in an ELISA and/or T-cell reactivity assay). Such immunogenic portions may react within such assays at a level that is similar to or greater than the reactivity of the full length polypeptide. Such screens may generally be performed using methods well known to those of ordinary skill in the art, such as those described in Harlow and Lane, *Antibodies: A Laboratory Manual*, Cold Spring Harbor Laboratory, 1988. For example, a polypeptide may be immobilized on a solid support and contacted with patient sera to allow binding of antibodies within the sera to the immobilized polypeptide. Unbound sera may then be removed and bound antibodies detected using, for example, ¹²⁵I-labeled Protein A.

As noted above, a composition may comprise a variant of a native lung tumor protein. A polypeptide "variant," as used herein, is a polypeptide that differs from a native lung tumor protein in one or more substitutions, deletions, additions and/or insertions, such that the immunogenicity of the polypeptide is not substantially diminished. In other words, the ability of a variant to react with antigen-specific antisera may be enhanced or unchanged, relative to the native protein, or may be diminished by less than 50%, and preferably less than 20%, relative to the native protein. Such variants may generally be identified by modifying one of the above polypeptide sequences and evaluating the reactivity of the modified polypeptide with antigen-specific antibodies or antisera as described herein. Preferred variants include

those in which one or more portions, such as an N-terminal leader sequence or transmembrane domain, have been removed. Other preferred variants include variants in which a small portion (e.g., 1-30 amino acids, preferably 5-15 amino acids) has been removed from the N- and/or C-terminal of the mature protein.

5 Polypeptide variants preferably exhibit at least about 70%, more preferably at least about 90% and most preferably at least about 95% identity (determined as described above) to the identified polypeptides.

Preferably, a variant contains conservative substitutions. A "conservative substitution" is one in which an amino acid is substituted for another amino acid that has similar properties, such that one skilled in the art of peptide chemistry would expect the secondary structure and hydropathic nature of the polypeptide to be substantially unchanged. Amino acid substitutions may generally be made on the basis of similarity in polarity, charge, solubility, hydrophobicity, hydrophilicity and/or the amphipathic nature of the residues. For example, negatively charged amino acids include aspartic acid and glutamic acid; positively charged amino acids include lysine and arginine; and amino acids with uncharged polar head groups having similar hydrophilicity values include leucine, isoleucine and valine; glycine and alanine; asparagine and glutamine; and serine, threonine, phenylalanine and tyrosine. Other groups of amino acids that may represent conservative changes include: (1) ala, pro, gly, glu, asp, gln, asn, ser, thr; (2) cys, ser, tyr, thr; (3) val, ile, leu, met, ala, phe; (4) lys, arg, his; and (5) phe, tyr, trp, his. A variant may also, or alternatively, contain nonconservative changes. In a preferred embodiment, variant polypeptides differ from a native sequence by substitution, deletion or addition of five amino acids or fewer. Variants may also (or alternatively) be modified by, for example, the deletion or addition of amino acids that have minimal influence on the immunogenicity, secondary structure and hydropathic nature of the polypeptide.

As noted above, polypeptides may comprise a signal (or leader) sequence at the N-terminal end of the protein which co-translationally or post-translationally directs transfer of the protein. The polypeptide may also be conjugated to a linker or other sequence for ease of synthesis, purification or identification of the

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polypeptide (e.g., poly-His), or to enhance binding of the polypeptide to a solid support. For example, a polypeptide may be conjugated to an immunoglobulin Fc region.

Polypeptides may be prepared using any of a variety of well known techniques. Recombinant polypeptides encoded by DNA sequences as described above may be readily prepared from the DNA sequences using any of a variety of expression vectors known to those of ordinary skill in the art. Expression may be achieved in any appropriate host cell that has been transformed or transfected with an expression vector containing a DNA molecule that encodes a recombinant polypeptide. Suitable host cells include prokaryotes, yeast, higher eukaryotic and plant cells. Preferably, the host cells employed are *E. coli*, yeast or a mammalian cell line such as COS or CHO. Supernatants from suitable host/vector systems which secrete recombinant protein or polypeptide into culture media may be first concentrated using a commercially available filter. Following concentration, the concentrate may be applied to a suitable purification matrix such as an affinity matrix or an ion exchange resin. Finally, one or more reverse phase HPLC steps can be employed to further purify a recombinant polypeptide.

Portions and other variants having fewer than about 100 amino acids, and generally fewer than about 50 amino acids, may also be generated by synthetic means, using techniques well known to those of ordinary skill in the art. For example, such polypeptides may be synthesized using any of the commercially available solid-phase techniques, such as the Merrifield solid-phase synthesis method, where amino acids are sequentially added to a growing amino acid chain. See Merrifield, *J. Am. Chem. Soc.* 85:2149-2146, 1963. Equipment for automated synthesis of polypeptides is commercially available from suppliers such as Perkin Elmer/Applied BioSystems Division (Foster City, CA), and may be operated according to the manufacturer's instructions.

Within certain specific embodiments, a polypeptide may be a fusion protein that comprises multiple polypeptides as described herein, or that comprises at least one polypeptide as described herein and an unrelated sequence, such as a known tumor protein. A fusion partner may, for example, assist in providing T helper epitopes (an immunological fusion partner), preferably T helper epitopes recognized by humans,

or may assist in expressing the protein (an expression enhancer) at higher yields than the native recombinant protein. Certain preferred fusion partners are both immunological and expression enhancing fusion partners. Other fusion partners may be selected so as to increase the solubility of the protein or to enable the protein to be targeted to desired intracellular compartments. Still further fusion partners include affinity tags, which facilitate purification of the protein.

Fusion proteins may generally be prepared using standard techniques, including chemical conjugation. Preferably, a fusion protein is expressed as a recombinant protein, allowing the production of increased levels, relative to a non-fused protein, in an expression system. Briefly, DNA sequences encoding the polypeptide components may be assembled separately, and ligated into an appropriate expression vector. The 3' end of the DNA sequence encoding one polypeptide component is ligated, with or without a peptide linker, to the 5' end of a DNA sequence encoding the second polypeptide component so that the reading frames of the sequences are in phase. This permits translation into a single fusion protein that retains the biological activity of both component polypeptides.

A peptide linker sequence may be employed to separate the first and the second polypeptide components by a distance sufficient to ensure that each polypeptide folds into its secondary and tertiary structures. Such a peptide linker sequence is incorporated into the fusion protein using standard techniques well known in the art. Suitable peptide linker sequences may be chosen based on the following factors: (1) their ability to adopt a flexible extended conformation; (2) their inability to adopt a secondary structure that could interact with functional epitopes on the first and second polypeptides; and (3) the lack of hydrophobic or charged residues that might react with the polypeptide functional epitopes. Preferred peptide linker sequences contain Gly, Asn and Ser residues. Other near neutral amino acids, such as Thr and Ala may also be used in the linker sequence. Amino acid sequences which may be usefully employed as linkers include those disclosed in Maratea et al., *Gene* 40:39-46, 1985; Murphy et al., *Proc. Natl. Acad. Sci. USA* 83:8258-8262, 1986; U.S. Patent No. 4,935,233 and U.S. Patent No. 4,751,180. The linker sequence may generally be from 1 to about 50 amino acids in length. Linker sequences are not required when the first and second

polypeptides have non-essential N-terminal amino acid regions that can be used to separate the functional domains and prevent steric interference.

The ligated DNA sequences are operably linked to suitable transcriptional or translational regulatory elements. The regulatory elements responsible for expression of DNA are located only 5' to the DNA sequence encoding the first polypeptides. Similarly, stop codons required to end translation and transcription termination signals are only present 3' to the DNA sequence encoding the second polypeptide.

Fusion proteins are also provided that comprise a polypeptide of the present invention together with an unrelated immunogenic protein. Preferably the immunogenic protein is capable of eliciting a recall response. Examples of such proteins include tetanus, tuberculosis and hepatitis proteins (*see, for example, Stoute et al. New Engl. J. Med.*, 336:86-91, 1997).

Within preferred embodiments, an immunological fusion partner is derived from protein D, a surface protein of the gram-negative bacterium *Haemophilus influenza B* (WO 91/18926). Preferably, a protein D derivative comprises approximately the first third of the protein (e.g., the first N-terminal 100-110 amino acids), and a protein D derivative may be lipidated. Within certain preferred embodiments, the first 109 residues of a Lipoprotein D fusion partner is included on the N-terminus to provide the polypeptide with additional exogenous T-cell epitopes and to increase the expression level in *E. coli* (thus functioning as an expression enhancer). The lipid tail ensures optimal presentation of the antigen to antigen presenting cells. Other fusion partners include the non-structural protein from influenzae virus, NS1 (hemagglutinin). Typically, the N-terminal 81 amino acids are used, although different fragments that include T-helper epitopes may be used.

In another embodiment, the immunological fusion partner is the protein known as LYTA, or a portion thereof (preferably a C-terminal portion). LYTA is derived from *Streptococcus pneumoniae*, which synthesizes an N-acetyl-L-alanine amidase known as amidase LYTA (encoded by the *LytA* gene; *Gene* 43:265-292, 1986). LYTA is an autolysin that specifically degrades certain bonds in the peptidoglycan backbone. The C-terminal domain of the LYTA protein is responsible

for the affinity to the choline or to some choline analogues such as DEAE. This property has been exploited for the development of *E. coli* C-LYTA expressing plasmids useful for expression of fusion proteins. Purification of hybrid proteins containing the C-LYTA fragment at the amino terminus has been described (see 5 *Biotechnology* 10:795-798, 1992). Within a preferred embodiment, a repeat portion of LYTA may be incorporated into a fusion protein. A repeat portion is found in the C-terminal region starting at residue 178. A particularly preferred repeat portion incorporates residues 188-305.

In general, polypeptides (including fusion proteins) and polynucleotides 10 as described herein are isolated. An "isolated" polypeptide or polynucleotide is one that is removed from its original environment. For example, a naturally-occurring protein is isolated if it is separated from some or all of the coexisting materials in the natural system. Preferably, such polypeptides are at least about 90% pure, more preferably at least about 95% pure and most preferably at least about 99% pure. A polynucleotide is 15 considered to be isolated if, for example, it is cloned into a vector that is not a part of the natural environment.

BINDING AGENTS

The present invention further provides agents, such as antibodies and 20 antigen-binding fragments thereof, that specifically bind to a lung tumor protein. As used herein, an antibody, or antigen-binding fragment thereof, is said to "specifically bind" to a lung tumor protein if it reacts at a detectable level (within, for example, an ELISA) with a lung tumor protein, and does not react detectably with unrelated proteins under similar conditions. As used herein, "binding" refers to a noncovalent association 25 between two separate molecules such that a complex is formed. The ability to bind may be evaluated by, for example, determining a binding constant for the formation of the complex. The binding constant is the value obtained when the concentration of the complex is divided by the product of the component concentrations. In general, two compounds are said to "bind," in the context of the present invention, when the binding 30 constant for complex formation exceeds about 10^3 L/mol. The binding constant may be determined using methods well known in the art.

Binding agents may be further capable of differentiating between patients with and without a cancer, such as lung cancer, using the representative assays provided herein. In other words, antibodies or other binding agents that bind to a lung tumor protein will generate a signal indicating the presence of a cancer in at least about 20% of patients with the disease, and will generate a negative signal indicating the absence of the disease in at least about 90% of individuals without the cancer. To determine whether a binding agent satisfies this requirement, biological samples (e.g., blood, sera, sputum urine and/or tumor biopsies) from patients with and without a cancer (as determined using standard clinical tests) may be assayed as described herein for the presence of polypeptides that bind to the binding agent. It will be apparent that a statistically significant number of samples with and without the disease should be assayed. Each binding agent should satisfy the above criteria; however, those of ordinary skill in the art will recognize that binding agents may be used in combination to improve sensitivity.

Any agent that satisfies the above requirements may be a binding agent. For example, a binding agent may be a ribosome, with or without a peptide component, an RNA molecule or a polypeptide. In a preferred embodiment, a binding agent is an antibody or an antigen-binding fragment thereof. Antibodies may be prepared by any of a variety of techniques known to those of ordinary skill in the art. See, e.g., Harlow and Lane, *Antibodies: A Laboratory Manual*, Cold Spring Harbor Laboratory, 1988. In general, antibodies can be produced by cell culture techniques, including the generation of monoclonal antibodies as described herein, or via transfection of antibody genes into suitable bacterial or mammalian cell hosts, in order to allow for the production of recombinant antibodies. In one technique, an immunogen comprising the polypeptide is initially injected into any of a wide variety of mammals (e.g., mice, rats, rabbits, sheep or goats). In this step, the polypeptides of this invention may serve as the immunogen without modification. Alternatively, particularly for relatively short polypeptides, a superior immune response may be elicited if the polypeptide is joined to a carrier protein, such as bovine serum albumin or keyhole limpet hemocyanin. The immunogen is injected into the animal host, preferably according to a predetermined schedule incorporating one or more booster immunizations, and the animals are bled periodically.

Polyclonal antibodies specific for the polypeptide may then be purified from such antisera by, for example, affinity chromatography using the polypeptide coupled to a suitable solid support.

Monoclonal antibodies specific for an antigenic polypeptide of interest may be prepared, for example, using the technique of Kohler and Milstein, *Eur. J. Immunol.* 6:511-519, 1976, and improvements thereto. Briefly, these methods involve the preparation of immortal cell lines capable of producing antibodies having the desired specificity (*i.e.*, reactivity with the polypeptide of interest). Such cell lines may be produced, for example, from spleen cells obtained from an animal immunized as described above. The spleen cells are then immortalized by, for example, fusion with a myeloma cell fusion partner, preferably one that is syngeneic with the immunized animal. A variety of fusion techniques may be employed. For example, the spleen cells and myeloma cells may be combined with a nonionic detergent for a few minutes and then plated at low density on a selective medium that supports the growth of hybrid cells, but not myeloma cells. A preferred selection technique uses HAT (hypoxanthine, aminopterin, thymidine) selection. After a sufficient time, usually about 1 to 2 weeks, colonies of hybrids are observed. Single colonies are selected and their culture supernatants tested for binding activity against the polypeptide. Hybridomas having high reactivity and specificity are preferred.

Monoclonal antibodies may be isolated from the supernatants of growing hybridoma colonies. In addition, various techniques may be employed to enhance the yield, such as injection of the hybridoma cell line into the peritoneal cavity of a suitable vertebrate host, such as a mouse. Monoclonal antibodies may then be harvested from the ascites fluid or the blood. Contaminants may be removed from the antibodies by conventional techniques, such as chromatography, gel filtration, precipitation, and extraction. The polypeptides of this invention may be used in the purification process in, for example, an affinity chromatography step.

Within certain embodiments, the use of antigen-binding fragments of antibodies may be preferred. Such fragments include Fab fragments, which may be prepared using standard techniques. Briefly, immunoglobulins may be purified from rabbit serum by affinity chromatography on Protein A bead columns (Harlow and Lane,

Antibodies: A Laboratory Manual, Cold Spring Harbor Laboratory, 1988) and digested by papain to yield Fab and Fc fragments. The Fab and Fc fragments may be separated by affinity chromatography on protein A bead columns.

Monoclonal antibodies of the present invention may be coupled to one or more therapeutic agents. Suitable agents in this regard include radionuclides, differentiation inducers, drugs, toxins, and derivatives thereof. Preferred radionuclides include ^{90}Y , ^{123}I , ^{125}I , ^{131}I , ^{186}Re , ^{188}Re , ^{211}At , and ^{212}Bi . Preferred drugs include methotrexate, and pyrimidine and purine analogs. Preferred differentiation inducers include phorbol esters and butyric acid. Preferred toxins include ricin, abrin, diphtheria toxin, cholera toxin, gelonin, *Pseudomonas* exotoxin, *Shigella* toxin, and pokeweed antiviral protein.

A therapeutic agent may be coupled (e.g., covalently bonded) to a suitable monoclonal antibody either directly or indirectly (e.g., via a linker group). A direct reaction between an agent and an antibody is possible when each possesses a substituent capable of reacting with the other. For example, a nucleophilic group, such as an amino or sulfhydryl group, on one may be capable of reacting with a carbonyl-containing group, such as an anhydride or an acid halide, or with an alkyl group containing a good leaving group (e.g., a halide) on the other.

Alternatively, it may be desirable to couple a therapeutic agent and an antibody via a linker group. A linker group can function as a spacer to distance an antibody from an agent in order to avoid interference with binding capabilities. A linker group can also serve to increase the chemical reactivity of a substituent on an agent or an antibody, and thus increase the coupling efficiency. An increase in chemical reactivity may also facilitate the use of agents, or functional groups on agents, which otherwise would not be possible.

It will be evident to those skilled in the art that a variety of bifunctional or polyfunctional reagents, both homo- and hetero-functional (such as those described in the catalog of the Pierce Chemical Co., Rockford, IL), may be employed as the linker group. Coupling may be effected, for example, through amino groups, carboxyl groups, sulfhydryl groups or oxidized carbohydrate residues. There are numerous references describing such methodology, e.g., U.S. Patent No. 4,671,958, to Rodwell et al.

Where a therapeutic agent is more potent when free from the antibody portion of the immunoconjugates of the present invention, it may be desirable to use a linker group which is cleavable during or upon internalization into a cell. A number of different cleavable linker groups have been described. The mechanisms for the intracellular release of an agent from these linker groups include cleavage by reduction of a disulfide bond (e.g., U.S. Patent No. 4,489,710, to Spitler), by irradiation of a photolabile bond (e.g., U.S. Patent No. 4,625,014, to Senter et al.), by hydrolysis of derivatized amino acid side chains (e.g., U.S. Patent No. 4,638,045, to Kohn et al.), by serum complement-mediated hydrolysis (e.g., U.S. Patent No. 4,671,958, to Rodwell et al.), and acid-catalyzed hydrolysis (e.g., U.S. Patent No. 4,569,789, to Blattler et al.).

It may be desirable to couple more than one agent to an antibody. In one embodiment, multiple molecules of an agent are coupled to one antibody molecule. In another embodiment, more than one type of agent may be coupled to one antibody. Regardless of the particular embodiment, immunoconjugates with more than one agent may be prepared in a variety of ways. For example, more than one agent may be coupled directly to an antibody molecule, or linkers which provide multiple sites for attachment can be used. Alternatively, a carrier can be used.

A carrier may bear the agents in a variety of ways, including covalent bonding either directly or via a linker group. Suitable carriers include proteins such as albumins (e.g., U.S. Patent No. 4,507,234, to Kato et al.), peptides and polysaccharides such as aminodextran (e.g., U.S. Patent No. 4,699,784, to Shih et al.). A carrier may also bear an agent by noncovalent bonding or by encapsulation, such as within a liposome vesicle (e.g., U.S. Patent Nos. 4,429,008 and 4,873,088). Carriers specific for radionuclide agents include radiohalogenated small molecules and chelating compounds. For example, U.S. Patent No. 4,735,792 discloses representative radiohalogenated small molecules and their synthesis. A radionuclide chelate may be formed from chelating compounds that include those containing nitrogen and sulfur atoms as the donor atoms for binding the metal, or metal oxide, radionuclide. For example, U.S. Patent No. 4,673,562, to Davison et al. discloses representative chelating compounds and their synthesis.

A variety of routes of administration for the antibodies and immunoconjugates may be used. Typically, administration will be intravenous, intramuscular, subcutaneous or in the bed of a resected tumor. It will be evident that the precise dose of the antibody/immunoconjugate will vary depending upon the antibody used, the antigen density on the tumor, and the rate of clearance of the antibody.

T CELLS

Immunotherapeutic compositions may also, or alternatively, comprise T cells specific for a lung tumor protein. Such cells may generally be prepared *in vitro* or *ex vivo*, using standard procedures. For example, T cells may be isolated from bone marrow, peripheral blood, or a fraction of bone marrow or peripheral blood of a patient, using a commercially available cell separation system, such as the Isolex™ System, available from Nexell Therapeutics, Inc. Irvine, CA (see also U.S. Patent No. 5,240,856; U.S. Patent No. 5,215,926; WO 89/06280; WO 91/16116 and WO 92/07243). Alternatively, T cells may be derived from related or unrelated humans, non-human mammals, cell lines or cultures.

T cells may be stimulated with a lung tumor polypeptide, polynucleotide encoding a lung tumor polypeptide and/or an antigen presenting cell (APC) that expresses such a polypeptide. Such stimulation is performed under conditions and for a time sufficient to permit the generation of T cells that are specific for the polypeptide. Preferably, a lung tumor polypeptide or polynucleotide is present within a delivery vehicle, such as a microsphere, to facilitate the generation of specific T cells.

T cells are considered to be specific for a lung tumor polypeptide if the T cells specifically proliferate, secrete cytokines or kill target cells coated with the polypeptide or expressing a gene encoding the polypeptide. T cell specificity may be evaluated using any of a variety of standard techniques. For example, within a chromium release assay or proliferation assay, a stimulation index of more than two fold increase in lysis and/or proliferation, compared to negative controls, indicates T cell specificity. Such assays may be performed, for example, as described in Chen et al., *Cancer Res.* 54:1065-1070, 1994. Alternatively, detection of the proliferation of T cells may be accomplished by a variety of known techniques. For example, T cell

proliferation can be detected by measuring an increased rate of DNA synthesis (e.g., by pulse-labeling cultures of T cells with tritiated thymidine and measuring the amount of tritiated thymidine incorporated into DNA). Contact with a lung tumor polypeptide (100 ng/ml - 100 µg/ml, preferably 200 ng/ml - 25 µg/ml) for 3 - 7 days should result in at least a two fold increase in proliferation of the T cells. Contact as described above for 2-3 hours should result in activation of the T cells, as measured using standard cytokine assays in which a two fold increase in the level of cytokine release (e.g., TNF or IFN-γ) is indicative of T cell activation (see Coligan et al., Current Protocols in Immunology, vol. 1, Wiley Interscience (Greene 1998)). T cells that have been activated in response to a lung tumor polypeptide, polynucleotide or polypeptide-expressing APC may be CD4⁺ and/or CD8⁺. Lung tumor protein-specific T cells may be expanded using standard techniques. Within preferred embodiments, the T cells are derived from either a patient or a related, or unrelated, donor and are administered to the patient following stimulation and expansion.

For therapeutic purposes, CD4⁺ or CD8⁺ T cells that proliferate in response to a lung tumor polypeptide, polynucleotide or APC can be expanded in number either *in vitro* or *in vivo*. Proliferation of such T cells *in vitro* may be accomplished in a variety of ways. For example, the T cells can be re-exposed to a lung tumor polypeptide, or a short peptide corresponding to an immunogenic portion of such a polypeptide, with or without the addition of T cell growth factors, such as interleukin-2, and/or stimulator cells that synthesize a lung tumor polypeptide. Alternatively, one or more T cells that proliferate in the presence of a lung tumor protein can be expanded in number by cloning. Methods for cloning cells are well known in the art, and include limiting dilution.

PHARMACEUTICAL COMPOSITIONS AND VACCINES

Within certain aspects, polypeptides, polynucleotides, T cells and/or binding agents disclosed herein may be incorporated into pharmaceutical compositions or immunogenic compositions (i.e., vaccines). Pharmaceutical compositions comprise one or more such compounds and a physiologically acceptable carrier. Vaccines may comprise one or more such compounds and an immunostimulant. An immunostimulant

may be any substance that enhances or potentiates an immune response to an exogenous antigen. Examples of immunostimulants include adjuvants, biodegradable microspheres (e.g., polylactic galactide) and liposomes (into which the compound is incorporated; see e.g., Fullerton, U.S. Patent No. 4,235,877). Vaccine preparation is generally described in, for example, M.F. Powell and M.J. Newman, eds., "Vaccine Design (the subunit and adjuvant approach)," Plenum Press (NY, 1995).

Pharmaceutical compositions and vaccines within the scope of the present invention may also contain other compounds, which may be biologically active or inactive. For example, one or more immunogenic portions of other tumor antigens may be present, either incorporated into a fusion polypeptide or as a separate compound, within the composition or vaccine.

A pharmaceutical composition or vaccine may contain DNA encoding one or more of the polypeptides as described above, such that the polypeptide is generated *in situ*. As noted above, the DNA may be present within any of a variety of delivery systems known to those of ordinary skill in the art, including nucleic acid expression systems, bacteria and viral expression systems. Numerous gene delivery techniques are well known in the art, such as those described by Rolland, *Crit. Rev. Therap. Drug Carrier Systems* 15:143-198, 1998, and references cited therein.

Appropriate nucleic acid expression systems contain the necessary DNA sequences for expression in the patient (such as a suitable promoter and terminating signal). Bacterial delivery systems involve the administration of a bacterium (such as *Bacillus-Calmette-Guerrin*) that expresses an immunogenic portion of the polypeptide on its cell surface or secretes such an epitope. In a preferred embodiment, the DNA may be introduced using a viral expression system (e.g., vaccinia or other pox virus, retrovirus, or adenovirus), which may involve the use of a non-pathogenic (defective), replication competent virus.

Suitable systems are disclosed, for example, in Fisher-Hoch et al., *Proc. Natl. Acad. Sci. USA* 86:317-321, 1989; Flexner et al., *Ann. N.Y. Acad. Sci.* 569:86-103, 1989; Flexner et al., *Vaccine* 8:17-21, 1990; U.S. Patent Nos. 4,603,112, 4,769,330, and 5,017,487; WO 89/01973; U.S. Patent No. 4,777,127; GB 2,200,651; EP 0,345,242; WO 91/02805; Berkner, *Biotechniques* 6:616-627, 1988; Rosenfeld et al., *Science* 252:431-434, 1991; Kolls et al., *Proc. Natl. Acad. Sci. USA* 91:215-219, 1994; Kass-Eisler et al., *Proc. Natl.*

Acad. Sci. USA 90:11498-11502, 1993; Guzman et al., *Circulation* 88:2838-2848, 1993; and Guzman et al., *Cir. Res.* 73:1202-1207, 1993. Techniques for incorporating DNA into such expression systems are well known to those of ordinary skill in the art. The DNA may also be "naked," as described, for example, in Ulmer et al., *Science* 5 259:1745-1749, 1993 and reviewed by Cohen, *Science* 259:1691-1692, 1993. The uptake of naked DNA may be increased by coating the DNA onto biodegradable beads, which are efficiently transported into the cells.

While any suitable carrier known to those of ordinary skill in the art may be employed in the pharmaceutical compositions of this invention, the type of carrier 10 will vary depending on the mode of administration. Compositions of the present invention may be formulated for any appropriate manner of administration, including for example, topical, oral, nasal, intravenous, intracranial, intraperitoneal, subcutaneous or intramuscular administration. For parenteral administration, such as subcutaneous injection, the carrier preferably comprises water, saline, alcohol, a fat, a wax or a buffer. 15 For oral administration, any of the above carriers or a solid carrier, such as mannitol, lactose, starch, magnesium stearate, sodium saccharine, talcum, cellulose, glucose, sucrose, and magnesium carbonate, may be employed. Biodegradable microspheres (e.g., polylactate polyglycolate) may also be employed as carriers for the pharmaceutical compositions of this invention. Suitable biodegradable microspheres 20 are disclosed, for example, in U.S. Patent Nos. 4,897,268 and 5,075,109.

Such compositions may also comprise buffers (e.g., neutral buffered saline or phosphate buffered saline), carbohydrates (e.g., glucose, mannose, sucrose or dextrans), mannitol, proteins, polypeptides or amino acids such as glycine, antioxidants, chelating agents such as EDTA or glutathione, adjuvants (e.g., aluminum hydroxide) 25 and/or preservatives. Alternatively, compositions of the present invention may be formulated as a lyophilizate. Compounds may also be encapsulated within liposomes using well known technology.

Any of a variety of immunostimulants may be employed in the vaccines of this invention. For example, an adjuvant may be included. Most adjuvants contain a 30 substance designed to protect the antigen from rapid catabolism, such as aluminum hydroxide or mineral oil, and a stimulator of immune responses, such as lipid A,

Bordetella pertussis or *Mycobacterium tuberculosis* derived proteins. Suitable adjuvants are commercially available as, for example, Freund's Incomplete Adjuvant and Complete Adjuvant (Difco Laboratories, Detroit, MI); Merck Adjuvant 65 (Merck and Company, Inc., Rahway, NJ); AS-2 (SmithKline Beecham, Philadelphia, PA);
5 aluminum salts such as aluminum hydroxide gel (alum) or aluminum phosphate; salts of calcium, iron or zinc; an insoluble suspension of acylated tyrosine; acylated sugars; cationically or anionically derivatized polysaccharides; polyphosphazenes; biodegradable microspheres; monophosphoryl lipid A and quil A. Cytokines, such as GM-CSF or interleukin-2, -7, or -12, may also be used as adjuvants.

10 Within the vaccines provided herein, the adjuvant composition is preferably designed to induce an immune response predominantly of the Th1 type. High levels of Th1-type cytokines (e.g., IFN- γ , TNF α , IL-2 and IL-12) tend to favor the induction of cell mediated immune responses to an administered antigen. In contrast, high levels of Th2-type cytokines (e.g., IL-4, IL-5, IL-6 and IL-10) tend to favor the
15 induction of humoral immune responses. Following application of a vaccine as provided herein, a patient will support an immune response that includes Th1- and Th2-type responses. Within a preferred embodiment, in which a response is predominantly Th1-type, the level of Th1-type cytokines will increase to a greater extent than the level of Th2-type cytokines. The levels of these cytokines may be readily assessed using
20 standard assays. For a review of the families of cytokines, see Mosmann and Coffman, *Ann. Rev. Immunol.* 7:145-173, 1989.

Preferred adjuvants for use in eliciting a predominantly Th1-type response include, for example, a combination of monophosphoryl lipid A, preferably 3-de-O-acylated monophosphoryl lipid A (3D-MPL), together with an aluminum salt.
25 MPL adjuvants are available from Ribi ImmunoChem Research Inc. (Hamilton, MT) (see US Patent Nos. 4,436,727; 4,877,611; 4,866,034 and 4,912,094). CpG-containing oligonucleotides (in which the CpG dinucleotide is unmethylated) also induce a predominantly Th1 response. Such oligonucleotides are well known and are described, for example, in WO 96/02555 and WO 99/33488. Immunostimulatory DNA sequences
30 are also described, for example, by Sato et al., *Science* 273:352, 1996. Another preferred adjuvant is a saponin, preferably QS21 (Aquila Biopharmaceuticals Inc.,

Framingham, MA), which may be used alone or in combination with other adjuvants. For example, an enhanced system involves the combination of a monophosphoryl lipid A and saponin derivative, such as the combination of QS21 and 3D-MPL as described in WO 94/00153, or a less reactogenic composition where the QS21 is quenched with cholesterol, as described in WO 96/33739. Other preferred formulations comprises an oil-in-water emulsion and tocopherol. A particularly potent adjuvant formulation involving QS21, 3D-MPL and tocopherol in an oil-in-water emulsion is described in WO 95/17210.

Other preferred adjuvants include Montanide ISA 720 (Seppic, France), SAF (Chiron, California, United States), ISCOMS (CSL), MF-59 (Chiron), the SBAS series of adjuvants (e.g., SBAS-2 or SBAS-4, available from SmithKline Beecham, Rixensart, Belgium), Detox (Ribi ImmunoChem Research Inc., Hamilton, MT), RC-529 (Ribi ImmunoChem Research Inc., Hamilton, MT) and Aminoalkyl glucosaminide 4-phosphates (AGPs).

Any vaccine provided herein may be prepared using well known methods that result in a combination of antigen, immune response enhancer and a suitable carrier or excipient. The compositions described herein may be administered as part of a sustained release formulation (i.e., a formulation such as a capsule, sponge or gel (composed of polysaccharides, for example) that effects a slow release of compound following administration). Such formulations may generally be prepared using well known technology (see, e.g. Coombes et al., *Vaccine* 14:1429-1438, 1996) and administered by, for example, oral, rectal or subcutaneous implantation, or by implantation at the desired target site. Sustained-release formulations may contain a polypeptide, polynucleotide or antibody dispersed in a carrier matrix and/or contained within a reservoir surrounded by a rate controlling membrane.

Carriers for use within such formulations are biocompatible, and may also be biodegradable; preferably the formulation provides a relatively constant level of active component release. Such carriers include microparticles of poly(lactide-co-glycolide), as well as polyacrylate, latex, starch, cellulose and dextran. Other delayed-release carriers include supramolecular biovectors, which comprise a non-liquid hydrophilic core (e.g., a cross-linked polysaccharide or oligosaccharide) and, optionally,

an external layer comprising an amphiphilic compound, such as a phospholipid (see e.g., U.S. Patent No. 5,151,254 and PCT applications WO 94/20078, WO/94/23701 and WO 96/06638). The amount of active compound contained within a sustained release formulation depends upon the site of implantation, the rate and expected duration of release and the nature of the condition to be treated or prevented.

Any of a variety of delivery vehicles may be employed within pharmaceutical compositions and vaccines to facilitate production of an antigen-specific immune response that targets tumor cells. Delivery vehicles include antigen presenting cells (APCs), such as dendritic cells, macrophages, B cells, monocytes and other cells that may be engineered to be efficient APCs. Such cells may, but need not, be genetically modified to increase the capacity for presenting the antigen, to improve activation and/or maintenance of the T cell response, to have anti-tumor effects *per se* and/or to be immunologically compatible with the receiver (*i.e.*, matched HLA haplotype). APCs may generally be isolated from any of a variety of biological fluids and organs, including tumor and peritumoral tissues, and may be autologous, allogeneic, syngeneic or xenogeneic cells.

Certain preferred embodiments of the present invention use dendritic cells or progenitors thereof as antigen-presenting cells. Dendritic cells are highly potent APCs (Banchereau and Steinman, *Nature* 392:245-251, 1998) and have been shown to be effective as a physiological adjuvant for eliciting prophylactic or therapeutic antitumor immunity (see Timmerman and Levy, *Ann. Rev. Med.* 50:507-529, 1999). In general, dendritic cells may be identified based on their typical shape (stellate *in situ*, with marked cytoplasmic processes (dendrites) visible *in vitro*); their ability to take up, process and present antigens with high efficiency, and their ability to activate naïve T cell responses. Dendritic cells may, of course, be engineered to express specific cell-surface receptors or ligands that are not commonly found on dendritic cells *in vivo* or *ex vivo*, and such modified dendritic cells are contemplated by the present invention. As an alternative to dendritic cells, secreted vesicles antigen-loaded dendritic cells (called exosomes) may be used within a vaccine (see Zitvogel et al., *Nature Med.* 4:594-600, 1998).

Dendritic cells and progenitors may be obtained from peripheral blood,

bone marrow, tumor-infiltrating cells, peritumoral tissues-infiltrating cells, lymph nodes, spleen, skin, umbilical cord blood or any other suitable tissue or fluid. For example, dendritic cells may be differentiated *ex vivo* by adding a combination of cytokines such as GM-CSF, IL-4, IL-13 and/or TNF α to cultures of monocytes
5 harvested from peripheral blood. Alternatively, CD34 positive cells harvested from peripheral blood, umbilical cord blood or bone marrow may be differentiated into dendritic cells by adding to the culture medium combinations of GM-CSF, IL-3, TNF α , CD40 ligand, LPS, flt3 ligand and/or other compound(s) that induce differentiation, maturation and proliferation of dendritic cells.

10 Dendritic cells are conveniently categorized as "immature" and "mature" cells, which allows a simple way to discriminate between two well characterized phenotypes. However, this nomenclature should not be construed to exclude all possible intermediate stages of differentiation. Immature dendritic cells are characterized as APC with a high capacity for antigen uptake and processing, which
15 correlates with the high expression of Fc γ receptor and mannose receptor. The mature phenotype is typically characterized by a lower expression of these markers, but a high expression of cell surface molecules responsible for T cell activation such as class I and class II MHC, adhesion molecules (e.g., CD54 and CD11) and costimulatory molecules (e.g., CD40, CD80, CD86 and 4-1BB).

20 APCs may generally be transfected with a polynucleotide encoding a lung tumor protein (or portion or other variant thereof) such that the lung tumor polypeptide, or an immunogenic portion thereof, is expressed on the cell surface. Such transfection may take place *ex vivo*, and a composition or vaccine comprising such transfected cells may then be used for therapeutic purposes, as described herein.

25 Alternatively, a gene delivery vehicle that targets a dendritic or other antigen presenting cell may be administered to a patient, resulting in transfection that occurs *in vivo*. *In vivo* and *ex vivo* transfection of dendritic cells, for example, may generally be performed using any methods known in the art, such as those described in WO 97/24447, or the gene gun approach described by Mahvi et al., *Immunology and cell*
30 *Biology* 75:456-460, 1997. Antigen loading of dendritic cells may be achieved by incubating dendritic cells or progenitor cells with the lung tumor polypeptide, DNA

(naked or within a plasmid vector) or RNA; or with antigen-expressing recombinant bacterium or viruses (e.g., vaccinia, fowlpox, adenovirus or lentivirus vectors). Prior to loading, the polypeptide may be covalently conjugated to an immunological partner that provides T cell help (e.g., a carrier molecule). Alternatively, a dendritic cell may be pulsed with a non-conjugated immunological partner, separately or in the presence of the polypeptide.

Vaccines and pharmaceutical compositions may be presented in unit-dose or multi-dose containers, such as sealed ampoules or vials. Such containers are preferably hermetically sealed to preserve sterility of the formulation until use. In general, formulations may be stored as suspensions, solutions or emulsions in oily or aqueous vehicles. Alternatively, a vaccine or pharmaceutical composition may be stored in a freeze-dried condition requiring only the addition of a sterile liquid carrier immediately prior to use.

15 CANCER THERAPY

In further aspects of the present invention, the compositions described herein may be used for immunotherapy of cancer, such as lung cancer. Within such methods, pharmaceutical compositions and vaccines are typically administered to a patient. As used herein, a "patient" refers to any warm-blooded animal, preferably a human. A patient may or may not be afflicted with cancer. Accordingly, the above pharmaceutical compositions and vaccines may be used to prevent the development of a cancer or to treat a patient afflicted with a cancer. A cancer may be diagnosed using criteria generally accepted in the art, including the presence of a malignant tumor. Pharmaceutical compositions and vaccines may be administered either prior to or following surgical removal of primary tumors and/or treatment such as administration of radiotherapy or conventional chemotherapeutic drugs.

Within certain embodiments, immunotherapy may be active immunotherapy, in which treatment relies on the *in vivo* stimulation of the endogenous host immune system to react against tumors with the administration of immune response-modifying agents (such as polypeptides and polynucleotides disclosed herein).

Within other embodiments, immunotherapy may be passive immunotherapy, in which treatment involves the delivery of agents with established tumor-immune reactivity (such as effector cells or antibodies) that can directly or indirectly mediate antitumor effects and does not necessarily depend on an intact host immune system. Examples of effector cells include T cells as discussed above, T lymphocytes (such as CD8⁺ cytotoxic T lymphocytes and CD4⁺ T-helper tumor-infiltrating lymphocytes), killer cells (such as Natural Killer cells and lymphokine-activated killer cells), B cells and antigen-presenting cells (such as dendritic cells and macrophages) expressing a polypeptide provided herein. T cell receptors and antibody receptors specific for the polypeptides recited herein may be cloned, expressed and transferred into other vectors or effector cells for adoptive immunotherapy. The polypeptides provided herein may also be used to generate antibodies or anti-idiotypic antibodies (as described above and in U.S. Patent No. 4,918,164) for passive immunotherapy.

Effector cells may generally be obtained in sufficient quantities for adoptive immunotherapy by growth *in vitro*, as described herein. Culture conditions for expanding single antigen-specific effector cells to several billion in number with retention of antigen recognition *in vivo* are well known in the art. Such *in vitro* culture conditions typically use intermittent stimulation with antigen, often in the presence of cytokines (such as IL-2) and non-dividing feeder cells. As noted above, immunoreactive polypeptides as provided herein may be used to rapidly expand antigen-specific T cell cultures in order to generate a sufficient number of cells for immunotherapy. In particular, antigen-presenting cells, such as dendritic, macrophage, monocyte, fibroblast and/or B cells, may be pulsed with immunoreactive polypeptides or transfected with one or more polynucleotides using standard techniques well known in the art. For example, antigen-presenting cells can be transfected with a polynucleotide having a promoter appropriate for increasing expression in a recombinant virus or other expression system. Cultured effector cells for use in therapy must be able to grow and distribute widely, and to survive long term *in vivo*. Studies have shown that cultured effector cells can be induced to grow *in vivo* and to survive

long term in substantial numbers by repeated stimulation with antigen supplemented with IL-2 (*see, for example, Cheever et al., Immunological Reviews 157:177, 1997*).

Alternatively, a vector expressing a polypeptide recited herein may be introduced into antigen presenting cells taken from a patient and clonally propagated *ex vivo* for transplant back into the same patient. Transfected cells may be reintroduced into the patient using any means known in the art, preferably in sterile form by intravenous, intracavitary, intraperitoneal or intratumor administration.

Routes and frequency of administration of the therapeutic compositions disclosed herein, as well as dosage, will vary from individual to individual, and may be readily established using standard techniques. In general, the pharmaceutical compositions and vaccines may be administered by injection (*e.g.*, intracutaneous, intramuscular, intravenous or subcutaneous), intranasally (*e.g.*, by aspiration) or orally. Preferably, between 1 and 10 doses may be administered over a 52 week period. Preferably, 6 doses are administered, at intervals of 1 month, and booster vaccinations may be given periodically thereafter. Alternate protocols may be appropriate for individual patients. A suitable dose is an amount of a compound that, when administered as described above, is capable of promoting an anti-tumor immune response, and is at least 10-50% above the basal (*i.e.*, untreated) level. Such response can be monitored by measuring the anti-tumor antibodies in a patient or by vaccine-dependent generation of cytolytic effector cells capable of killing the patient's tumor cells *in vitro*. Such vaccines should also be capable of causing an immune response that leads to an improved clinical outcome (*e.g.*, more frequent remissions, complete or partial or longer disease-free survival) in vaccinated patients as compared to non-vaccinated patients. In general, for pharmaceutical compositions and vaccines comprising one or more polypeptides, the amount of each polypeptide present in a dose ranges from about 25 μ g to 5 mg per kg of host. Suitable dose sizes will vary with the size of the patient, but will typically range from about 0.1 mL to about 5 mL.

In general, an appropriate dosage and treatment regimen provides the active compound(s) in an amount sufficient to provide therapeutic and/or prophylactic benefit. Such a response can be monitored by establishing an improved clinical outcome (*e.g.*, more frequent remissions, complete or partial, or longer disease-free

survival) in treated patients as compared to non-treated patients. Increases in preexisting immune responses to a lung tumor protein generally correlate with an improved clinical outcome. Such immune responses may generally be evaluated using standard proliferation, cytotoxicity or cytokine assays, which may be performed using samples obtained from a patient before and after treatment.

METHODS FOR DETECTING CANCER

In general, a cancer may be detected in a patient based on the presence of one or more lung tumor proteins and/or polynucleotides encoding such proteins in a biological sample (for example, blood, sera, sputum urine and/or tumor biopsies) obtained from the patient. In other words, such proteins may be used as markers to indicate the presence or absence of a cancer such as lung cancer. In addition, such proteins may be useful for the detection of other cancers. The binding agents provided herein generally permit detection of the level of antigen that binds to the agent in the biological sample. Polynucleotide primers and probes may be used to detect the level of mRNA encoding a tumor protein, which is also indicative of the presence or absence of a cancer. In general, a lung tumor sequence should be present at a level that is at least three fold higher in tumor tissue than in normal tissue.

There are a variety of assay formats known to those of ordinary skill in the art for using a binding agent to detect polypeptide markers in a sample. See, e.g., Harlow and Lane, *Antibodies: A Laboratory Manual*, Cold Spring Harbor Laboratory, 1988. In general, the presence or absence of a cancer in a patient may be determined by (a) contacting a biological sample obtained from a patient with a binding agent; (b) detecting in the sample a level of polypeptide that binds to the binding agent; and (c) comparing the level of polypeptide with a predetermined cut-off value.

In a preferred embodiment, the assay involves the use of binding agent immobilized on a solid support to bind to and remove the polypeptide from the remainder of the sample. The bound polypeptide may then be detected using a detection reagent that contains a reporter group and specifically binds to the binding agent/polypeptide complex. Such detection reagents may comprise, for example, a binding agent that specifically binds to the polypeptide or an antibody or other agent

that specifically binds to the binding agent, such as an anti-immunoglobulin, protein G, protein A or a lectin. Alternatively, a competitive assay may be utilized, in which a polypeptide is labeled with a reporter group and allowed to bind to the immobilized binding agent after incubation of the binding agent with the sample. The extent to which components of the sample inhibit the binding of the labeled polypeptide to the binding agent is indicative of the reactivity of the sample with the immobilized binding agent. Suitable polypeptides for use within such assays include full length lung tumor proteins and portions thereof to which the binding agent binds, as described above.

The solid support may be any material known to those of ordinary skill in the art to which the tumor protein may be attached. For example, the solid support may be a test well in a microtiter plate or a nitrocellulose or other suitable membrane. Alternatively, the support may be a bead or disc, such as glass, fiberglass, latex or a plastic material such as polystyrene or polyvinylchloride. The support may also be a magnetic particle or a fiber optic sensor, such as those disclosed, for example, in U.S. Patent No. 5,359,681. The binding agent may be immobilized on the solid support using a variety of techniques known to those of skill in the art, which are amply described in the patent and scientific literature. In the context of the present invention, the term "immobilization" refers to both noncovalent association, such as adsorption, and covalent attachment (which may be a direct linkage between the agent and functional groups on the support or may be a linkage by way of a cross-linking agent). Immobilization by adsorption to a well in a microtiter plate or to a membrane is preferred. In such cases, adsorption may be achieved by contacting the binding agent, in a suitable buffer, with the solid support for a suitable amount of time. The contact time varies with temperature, but is typically between about 1 hour and about 1 day. In general, contacting a well of a plastic microtiter plate (such as polystyrene or polyvinylchloride) with an amount of binding agent ranging from about 10 ng to about 10 μ g, and preferably about 100 ng to about 1 μ g, is sufficient to immobilize an adequate amount of binding agent.

Covalent attachment of binding agent to a solid support may generally be achieved by first reacting the support with a bifunctional reagent that will react with both the support and a functional group, such as a hydroxyl or amino group, on the

binding agent. For example, the binding agent may be covalently attached to supports having an appropriate polymer coating using benzoquinone or by condensation of an aldehyde group on the support with an amine and an active hydrogen on the binding partner (see, e.g., Pierce Immunotechnology Catalog and Handbook, 1991, at 5 A12-A13).

In certain embodiments, the assay is a two-antibody sandwich assay. This assay may be performed by first contacting an antibody that has been immobilized on a solid support, commonly the well of a microtiter plate, with the sample, such that polypeptides within the sample are allowed to bind to the immobilized antibody. 10 Unbound sample is then removed from the immobilized polypeptide-antibody complexes and a detection reagent (preferably a second antibody capable of binding to a different site on the polypeptide) containing a reporter group is added. The amount of detection reagent that remains bound to the solid support is then determined using a method appropriate for the specific reporter group.

15 More specifically, once the antibody is immobilized on the support as described above, the remaining protein binding sites on the support are typically blocked. Any suitable blocking agent known to those of ordinary skill in the art, such as bovine serum albumin or Tween 20™ (Sigma Chemical Co., St. Louis, MO). The immobilized antibody is then incubated with the sample, and polypeptide is allowed to 20 bind to the antibody. The sample may be diluted with a suitable diluent, such as phosphate-buffered saline (PBS) prior to incubation. In general, an appropriate contact time (i.e., incubation time) is a period of time that is sufficient to detect the presence of polypeptide within a sample obtained from an individual with lung cancer. Preferably, the contact time is sufficient to achieve a level of binding that is at least about 95% of 25 that achieved at equilibrium between bound and unbound polypeptide. Those of ordinary skill in the art will recognize that the time necessary to achieve equilibrium may be readily determined by assaying the level of binding that occurs over a period of time. At room temperature, an incubation time of about 30 minutes is generally sufficient.

30 Unbound sample may then be removed by washing the solid support with an appropriate buffer, such as PBS containing 0.1% Tween 20™. The second

antibody, which contains a reporter group, may then be added to the solid support. Preferred reporter groups include those groups recited above.

The detection reagent is then incubated with the immobilized antibody-polypeptide complex for an amount of time sufficient to detect the bound polypeptide.

5 An appropriate amount of time may generally be determined by assaying the level of binding that occurs over a period of time. Unbound detection reagent is then removed and bound detection reagent is detected using the reporter group. The method employed for detecting the reporter group depends upon the nature of the reporter group. For radioactive groups, scintillation counting or autoradiographic methods are
10 generally appropriate. Spectroscopic methods may be used to detect dyes, luminescent groups and fluorescent groups. Biotin may be detected using avidin, coupled to a different reporter group (commonly a radioactive or fluorescent group or an enzyme). Enzyme reporter groups may generally be detected by the addition of substrate (generally for a specific period of time), followed by spectroscopic or other analysis of
15 the reaction products.

To determine the presence or absence of a cancer, such as lung cancer, the signal detected from the reporter group that remains bound to the solid support is generally compared to a signal that corresponds to a predetermined cut-off value. In one preferred embodiment, the cut-off value for the detection of a cancer is the average
20 mean signal obtained when the immobilized antibody is incubated with samples from patients without the cancer. In general, a sample generating a signal that is three standard deviations above the predetermined cut-off value is considered positive for the cancer. In an alternate preferred embodiment, the cut-off value is determined using a Receiver Operator Curve, according to the method of Sackett et al., *Clinical*
25 *Epidemiology: A Basic Science for Clinical Medicine*, Little Brown and Co., 1985, p. 106-7. Briefly, in this embodiment, the cut-off value may be determined from a plot of pairs of true positive rates (i.e., sensitivity) and false positive rates (100%-specificity) that correspond to each possible cut-off value for the diagnostic test result. The cut-off value on the plot that is the closest to the upper left-hand corner (i.e., the value that
30 encloses the largest area) is the most accurate cut-off value, and a sample generating a signal that is higher than the cut-off value determined by this method may be considered

positive. Alternatively, the cut-off value may be shifted to the left along the plot, to minimize the false positive rate, or to the right, to minimize the false negative rate. In general, a sample generating a signal that is higher than the cut-off value determined by this method is considered positive for a cancer.

5 In a related embodiment, the assay is performed in a flow-through or strip test format, wherein the binding agent is immobilized on a membrane, such as nitrocellulose. In the flow-through test, polypeptides within the sample bind to the immobilized binding agent as the sample passes through the membrane. A second, labeled binding agent then binds to the binding agent-polypeptide complex as a solution
10 containing the second binding agent flows through the membrane. The detection of bound second binding agent may then be performed as described above. In the strip test format, one end of the membrane to which binding agent is bound is immersed in a solution containing the sample. The sample migrates along the membrane through a region containing second binding agent and to the area of immobilized binding agent.
15 Concentration of second binding agent at the area of immobilized antibody indicates the presence of a cancer. Typically, the concentration of second binding agent at that site generates a pattern, such as a line, that can be read visually. The absence of such a pattern indicates a negative result. In general, the amount of binding agent immobilized on the membrane is selected to generate a visually discernible pattern when the
20 biological sample contains a level of polypeptide that would be sufficient to generate a positive signal in the two-antibody sandwich assay, in the format discussed above. Preferred binding agents for use in such assays are antibodies and antigen-binding fragments thereof. Preferably, the amount of antibody immobilized on the membrane ranges from about 25 ng to about 1 µg, and more preferably from about 50 ng to about
25 500 ng. Such tests can typically be performed with a very small amount of biological sample.

Of course, numerous other assay protocols exist that are suitable for use with the tumor proteins or binding agents of the present invention. The above descriptions are intended to be exemplary only. For example, it will be apparent to
30 those of ordinary skill in the art that the above protocols may be readily modified to use lung tumor polypeptides to detect antibodies that bind to such polypeptides in a

biological sample. The detection of such lung tumor protein specific antibodies may correlate with the presence of a cancer.

A cancer may also, or alternatively, be detected based on the presence of T cells that specifically react with a lung tumor protein in a biological sample. Within certain methods, a biological sample comprising CD4⁺ and/or CD8⁺ T cells isolated from a patient is incubated with a lung tumor polypeptide, a polynucleotide encoding such a polypeptide and/or an APC that expresses at least an immunogenic portion of such a polypeptide, and the presence or absence of specific activation of the T cells is detected. Suitable biological samples include, but are not limited to, isolated T cells. For example, T cells may be isolated from a patient by routine techniques (such as by Ficoll/Hypaque density gradient centrifugation of peripheral blood lymphocytes). T cells may be incubated *in vitro* for 2-9 days (typically 4 days) at 37°C with polypeptide (e.g., 5 - 25 µg/ml). It may be desirable to incubate another aliquot of a T cell sample in the absence of lung tumor polypeptide to serve as a control. For CD4⁺ T cells, activation is preferably detected by evaluating proliferation of the T cells. For CD8⁺ T cells, activation is preferably detected by evaluating cytolytic activity. A level of proliferation that is at least two fold greater and/or a level of cytolytic activity that is at least 20% greater than in disease-free patients indicates the presence of a cancer in the patient.

As noted above, a cancer may also, or alternatively, be detected based on the level of mRNA encoding a lung tumor protein in a biological sample. For example, at least two oligonucleotide primers may be employed in a polymerase chain reaction (PCR) based assay to amplify a portion of a lung tumor cDNA derived from a biological sample, wherein at least one of the oligonucleotide primers is specific for (i.e., hybridizes to) a polynucleotide encoding the lung tumor protein. The amplified cDNA is then separated and detected using techniques well known in the art, such as gel electrophoresis. Similarly, oligonucleotide probes that specifically hybridize to a polynucleotide encoding a lung tumor protein may be used in a hybridization assay to detect the presence of polynucleotide encoding the tumor protein in a biological sample.

To permit hybridization under assay conditions, oligonucleotide primers and probes should comprise an oligonucleotide sequence that has at least about 60%,

preferably at least about 75% and more preferably at least about 90%, identity to a portion of a polynucleotide encoding a lung tumor protein that is at least 10 nucleotides, and preferably at least 20 nucleotides, in length. Preferably, oligonucleotide primers and/or probes will hybridize to a polynucleotide encoding a polypeptide disclosed herein under moderately stringent conditions, as defined above. Oligonucleotide primers and/or probes which may be usefully employed in the diagnostic methods described herein preferably are at least 10-40 nucleotides in length. In a preferred embodiment, the oligonucleotide primers comprise at least 10 contiguous nucleotides, more preferably at least 15 contiguous nucleotides, of a DNA molecule having a sequence recited in SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349. Techniques for both PCR based assays and hybridization assays are well known in the art (*see*, for example, Mullis et al., *Cold Spring Harbor Symp. Quant. Biol.*, 51:263, 1987; Erlich ed., *PCR Technology*, Stockton Press, NY, 1989).

One preferred assay employs RT-PCR, in which PCR is applied in conjunction with reverse transcription. Typically, RNA is extracted from a biological sample, such as biopsy tissue, and is reverse transcribed to produce cDNA molecules. PCR amplification using at least one specific primer generates a cDNA molecule, which may be separated and visualized using, for example, gel electrophoresis. Amplification may be performed on biological samples taken from a test patient and from an individual who is not afflicted with a cancer. The amplification reaction may be performed on several dilutions of cDNA spanning two orders of magnitude. A two-fold or greater increase in expression in several dilutions of the test patient sample as compared to the same dilutions of the non-cancerous sample is typically considered positive.

In another embodiment, the disclosed compositions may be used as markers for the progression of cancer. In this embodiment, assays as described above for the diagnosis of a cancer may be performed over time, and the change in the level of reactive polypeptide(s) or polynucleotide evaluated. For example, the assays may be performed every 24-72 hours for a period of 6 months to 1 year, and thereafter performed as needed. In general, a cancer is progressing in those patients in whom the

level of polypeptide or polynucleotide detected increases over time. In contrast, the cancer is not progressing when the level of reactive polypeptide or polynucleotide either remains constant or decreases with time.

Certain *in vivo* diagnostic assays may be performed directly on a tumor.

- 5 One such assay involves contacting tumor cells with a binding agent. The bound binding agent may then be detected directly or indirectly via a reporter group. Such binding agents may also be used in histological applications. Alternatively, polynucleotide probes may be used within such applications.

- As noted above, to improve sensitivity, multiple lung tumor protein
10 markers may be assayed within a given sample. It will be apparent that binding agents specific for different proteins provided herein may be combined within a single assay. Further, multiple primers or probes may be used concurrently. The selection of tumor protein markers may be based on routine experiments to determine combinations that results in optimal sensitivity. In addition, or alternatively, assays for tumor proteins
15 provided herein may be combined with assays for other known tumor antigens.

DIAGNOSTIC KITS

- The present invention further provides kits for use within any of the above diagnostic methods. Such kits typically comprise two or more components
20 necessary for performing a diagnostic assay. Components may be compounds, reagents, containers and/or equipment. For example, one container within a kit may contain a monoclonal antibody or fragment thereof that specifically binds to a lung tumor protein. Such antibodies or fragments may be provided attached to a support material, as described above. One or more additional containers may enclose elements,
25 such as reagents or buffers, to be used in the assay. Such kits may also, or alternatively, contain a detection reagent as described above that contains a reporter group suitable for direct or indirect detection of antibody binding.

- Alternatively, a kit may be designed to detect the level of mRNA encoding a lung tumor protein in a biological sample. Such kits generally comprise at
30 least one oligonucleotide probe or primer, as described above, that hybridizes to a polynucleotide encoding a lung tumor protein. Such an oligonucleotide may be used,

for example, within a PCR or hybridization assay. Additional components that may be present within such kits include a second oligonucleotide and/or a diagnostic reagent or container to facilitate the detection of a polynucleotide encoding a lung tumor protein.

The following Examples are offered by way of illustration and not by way of limitation.

EXAMPLE 1

ISOLATION AND CHARACTERIZATION OF cDNA SEQUENCES
ENCODING LUNG TUMOR POLYPEPTIDES

5 This example illustrates the isolation of cDNA molecules encoding lung tumor-specific polypeptides from lung tumor cDNA libraries.

A. ISOLATION OF cDNA SEQUENCES FROM A LUNG SQUAMOUS CELL
10 CARCINOMA LIBRARY

A human lung squamous cell carcinoma cDNA expression library was constructed from poly A⁺ RNA from a pool of two patient tissues using a Superscript Plasmid System for cDNA Synthesis and Plasmid Cloning kit (BRL Life Technologies, Gaithersburg, MD) following the manufacturer's protocol. Specifically, lung carcinoma
15 tissues were homogenized with polytron (Kinematica, Switzerland) and total RNA was extracted using Trizol reagent (BRL Life Technologies) as directed by the manufacturer. The poly A⁺ RNA was then purified using an oligo dT cellulose column as described in Sambrook et al., *Molecular Cloning: A Laboratory Manual*, Cold Spring Harbor Laboratories, Cold Spring Harbor, NY, 1989. First-strand cDNA was
20 synthesized using the NotI/Oligo-dT18 primer. Double-stranded cDNA was synthesized, ligated with BstXI/EcoRI adaptors (Invitrogen, San Diego, CA) and digested with NotI. Following size fractionation with cDNA size fractionation columns (BRL Life Technologies), the cDNA was ligated into the BstXI/NotI site of pcDNA3.1 (Invitrogen) and transformed into ElectroMax *E. coli* DH10B cells (BRL Life
25 Technologies) by electroporation.

Using the same procedure, a normal human lung cDNA expression library was prepared from a pool of four tissue specimens. The cDNA libraries were characterized by determining the number of independent colonies, the percentage of clones that carried insert, the average insert size and by sequence analysis. The lung
30 squamous cell carcinoma library contained 2.7×10^6 independent colonies, with 100% of clones having an insert and the average insert size being 2100 base pairs. The normal

lung cDNA library contained 1.4×10^6 independent colonies, with 90% of clones having inserts and the average insert size being 1800 base pairs. For both libraries, sequence analysis showed that the majority of clones had a full-length cDNA sequence and were synthesized from mRNA.

5 cDNA library subtraction was performed using the above lung squamous cell carcinoma and normal lung cDNA libraries, as described by Hara *et al.* (*Blood*, 84:189-199, 1994) with some modifications. Specifically, a lung squamous cell carcinoma-specific subtracted cDNA library was generated as follows. Normal tissue cDNA library (80 μ g) was digested with BamHI and XhoI, followed by a filling-in
10 reaction with DNA polymerase Klenow fragment. After phenol-chloroform extraction and ethanol precipitation, the DNA was dissolved in 133 μ l of H₂O, heat-denatured and mixed with 133 μ l (133 μ g) of Photoprobe biotin (Vector Laboratories, Burlingame, CA). As recommended by the manufacturer, the resulting mixture was irradiated with a 270 W sunlamp on ice for 20 minutes. Additional Photoprobe biotin (67 μ l) was added
15 and the biotinylation reaction was repeated. After extraction with butanol five times, the DNA was ethanol-precipitated and dissolved in 23 μ l H₂O to form the driver DNA.
To form the tracer DNA, 10 μ g lung squamous cell carcinoma cDNA library was digested with NotI and SpeI, phenol chloroform extracted and passed through Chroma spin-400 columns (Clontech, Palo Alto, CA). Typically, 5 μ g of
20 cDNA was recovered after the sizing column. Following ethanol precipitation, the tracer DNA was dissolved in 5 μ l H₂O. Tracer DNA was mixed with 15 μ l driver DNA and 20 μ l of 2 x hybridization buffer (1.5 M NaCl/10 mM EDTA/50 mM HEPES pH 7.5/0.2% sodium dodecyl sulfate), overlaid with mineral oil, and heat-denatured completely. The sample was immediately transferred into a 68 °C water bath and
25 incubated for 20 hours (long hybridization [LH]). The reaction mixture was then subjected to a streptavidin treatment followed by phenol/chloroform extraction. This process was repeated three more times. Subtracted DNA was precipitated, dissolved in 12 μ l H₂O, mixed with 8 μ l driver DNA and 20 μ l of 2 x hybridization buffer, and subjected to a hybridization at 68 °C for 2 hours (short hybridization [SH]). After
30 removal of biotinylated double-stranded DNA, subtracted cDNA was ligated into NotI/SpeI site of chloramphenicol resistant pBCSK⁺ (Stratagene, La Jolla, CA) and

transformed into ElectroMax *E. coli* DH10B cells by electroporation to generate a lung squamous cell carcinoma specific subtracted cDNA library (herein after referred to as "lung subtraction I").

A second lung squamous cell carcinoma specific subtracted cDNA library (referred to as "lung subtraction II") was generated in a similar way to the lung subtraction library I, except that eight frequently recovered genes from lung subtraction I were included in the driver DNA, and 24,000 independent clones were recovered.

To analyze the subtracted cDNA libraries, plasmid DNA was prepared from 320 independent clones, randomly picked from the subtracted lung squamous cell carcinoma specific libraries. Representative cDNA clones were further characterized by DNA sequencing with a Perkin Elmer/Applied Biosystems Division Automated Sequencer Model 373A and/or Model 377 (Foster City, CA). The cDNA sequences for sixty isolated clones are provided in SEQ ID NO: 1-60. These sequences were compared to known sequences in the gene bank using the EMBL and GenBank databases (release 96). No significant homologies were found to the sequences provided in SEQ ID NO: 2, 3, 19, 38 and 46. The sequences of SEQ ID NO: 1, 6-8, 10-13, 15, 17, 18, 20-27, 29, 30, 32, 34-37, 39-45, 47-49, 51, 52, 54, 55 and 57-59 were found to show some homology to previously identified expressed sequence tags (ESTs). The sequences of SEQ ID NO: 9, 28, 31 and 33 were found to show some homology to previously identified non-human gene sequences and the sequences of SEQ ID NO: 4, 5, 14, 50, 53, 56 and 60 were found to show some homology to gene sequences previously identified in humans.

The subtraction procedure described above was repeated using the above lung squamous cell carcinoma cDNA library as the tracer DNA, and the above normal lung tissue cDNA library and a cDNA library from normal liver and heart (constructed from a pool of one sample of each tissue as described above), plus twenty other cDNA clones that were frequently recovered in lung subtractions I and II, as the driver DNA (lung subtraction III). The normal liver and heart cDNA library contained 1.76×10^6 independent colonies, with 100% of clones having inserts and the average insert size being 1600 base pairs. Ten additional clones were isolated (SEQ ID NO: 61-70). Comparison of these cDNA sequences with those in the gene bank as described above,

revealed no significant homologies to the sequences provided in SEQ ID NO: 62 and 67. The sequences of SEQ ID NO: 61, 63-66, 68 and 69 were found to show some homology to previously isolated ESTs and the sequence provided in SEQ ID NO: 70 was found to show some homology to a previously identified rat gene.

5 In further studies, the subtraction procedure described above was repeated using the above lung squamous cell carcinoma cDNA library as the tracer DNA, and a cDNA library from a pool of normal lung, kidney, colon, pancreas, brain, resting PBMC, heart, skin and esophagus as the driver DNA, with esophagus cDNAs making up one third of the driver material. Since esophagus is enriched in normal
10 epithelial cells, including differentiated squamous cells, this procedure is likely to enrich genes that are tumor specific rather than tissues specific. The cDNA sequences of 48 clones determined in this subtraction are provided in SEQ ID NO: 177-224. The sequences of SEQ ID NO: 177, 178, 180, 181, 183, 187, 192, 195-197, 208, 211, 212, 215, 216, 218 and 219 showed some homology to previously identified genes. The
15 sequences of SEQ ID NO: 179, 182, 184-186, 188-191, 193, 194, 198-207, 209 210, 213, 214, 217, 220 and 224 showed some homology to previously determined ESTs. The sequence of SEQ ID NO: 221-223 showed no homology to any previously determined sequence.

20 B. ISOLATION OF cDNA SEQUENCES FROM A LUNG ADENOCARCINOMA LIBRARY

A human lung adenocarcinoma cDNA expression library was constructed as described above. The library contained 3.2×10^6 independent colonies, with 100% of clones having an insert and the average insert size being 1500 base pairs.
25 Library subtraction was performed as described above using the normal lung and normal liver and heart cDNA expression libraries described above as the driver DNA. Twenty-six hundred independent clones were recovered.

Initial cDNA sequence analysis from 100 independent clones revealed many ribosomal protein genes. The cDNA sequences for fifteen clones isolated in this
30 subtraction are provided in SEQ ID NO: 71-86. Comparison of these sequences with those in the gene bank as described above revealed no significant homologies to the

sequence provided in SEQ ID NO: 84. The sequences of SEQ ID NO: 71, 73, 74, 77, 78 and 80-82 were found to show some homology to previously isolated ESTs, and the sequences of SEQ ID NO: 72, 75, 76, 79, 83 and 85 were found to show some homology to previously identified human genes.

5 In further studies, a cDNA library (referred to as mets3616A) was constructed from a metastatic lung adenocarcinoma. The determined cDNA sequences of 25 clones sequenced at random from this library are provided in SEQ ID NO: 255-279. The mets3616A cDNA library was subtracted against a cDNA library prepared from a pool of normal lung, liver, pancreas, skin, kidney, brain and resting PBMC. To
10 increase the specificity of the subtraction, the driver was spiked with genes that were determined to be most abundant in the mets3616A cDNA library, such as EF1-alpha, integrin-beta and anticoagulant protein PP4, as well as with cDNAs that were previously found to be differentially expressed in subtracted lung adenocarcinoma cDNA libraries. The determined cDNA sequences of 51 clones isolated from the
15 subtracted library (referred to as mets3616A-S1) are provided in SEQ ID NO: 280-330.

Comparison of the sequences of SEQ ID NO: 255-330 with those in the public databases revealed no significant homologies to the sequences of SEQ ID NO: 255-258, 260, 262-264, 270, 272, 275, 276, 279, 281, 287, 291, 296, 300 and 310. The sequences of SEQ ID NO: 259, 261, 265-269, 271, 273, 274, 277, 278, 282-285, 288-
20 290, 292, 294, 297-299, 301, 303-309, 313, 314, 316, 320-324 and 326-330 showed some homology to previously identified gene sequences, while the sequences of SEQ ID NO: 280, 286, 293, 302, 310, 312, 315, 317-319 and 325 showed some homology to previously isolated expressed sequence tags (ESTs).

25

EXAMPLE 2

DETERMINATION OF TISSUE SPECIFICITY OF LUNG TUMOR

POLYPEPTIDES

Using gene specific primers, mRNA expression levels for seven
30 representative lung tumor polypeptides described in Example 1 were examined in a variety of normal and tumor tissues using RT-PCR.

Briefly, total RNA was extracted from a variety of normal and tumor tissues using Trizol reagent as described above. First strand synthesis was carried out using 2 μ g of total RNA with SuperScript II reverse transcriptase (BRL Life Technologies) at 42 °C for one hour. The cDNA was then amplified by PCR with gene-specific primers. To ensure the semi-quantitative nature of the RT-PCR, β -actin was used as an internal control for each of the tissues examined. 1 μ l of 1:30 dilution of cDNA was employed to enable the linear range amplification of the β -actin template and was sensitive enough to reflect the differences in the initial copy numbers. Using these conditions, the β -actin levels were determined for each reverse transcription reaction from each tissue. DNA contamination was minimized by DNase treatment and by assuring a negative PCR result when using first strand cDNA that was prepared without adding reverse transcriptase.

mRNA Expression levels were examined in five different types of tumor tissue (lung squamous cell carcinoma from 3 patients, lung adenocarcinoma, colon tumor from 2 patients, breast tumor and prostate tumor), and thirteen different normal tissues (lung from 4 donors, prostate, brain, kidney, liver, ovary, skeletal muscle, skin, small intestine, stomach, myocardium, retina and testes). Using a 10-fold amount of cDNA, the antigen LST-S1-90 (SEQ ID NO: 3) was found to be expressed at high levels in lung squamous cell carcinoma and in breast tumor, and at low to undetectable levels in the other tissues examined.

The antigen LST-S2-68 (SEQ ID NO: 15) appears to be specific to lung and breast tumor, however, expression was also detected in normal kidney. Antigens LST-S1-169 (SEQ ID NO: 6) and LST-S1-133 (SEQ ID NO: 5) appear to be very abundant in lung tissues (both normal and tumor), with the expression of these two genes being decreased in most of the normal tissues tested. Both LST-S1-169 and LST-S1-133 were also expressed in breast and colon tumors. Antigens LST-S1-6 (SEQ ID NO: 7) and LST-S2-I2-5F (SEQ ID NO: 47) did not show tumor or tissue specific expression, with the expression of LST-S1-28 being rare and only detectable in a few tissues. The antigen LST-S3-7 (SEQ ID NO: 63) showed lung and breast tumor specific expression, with its message only being detected in normal testes when the PCR was performed for 30 cycles. Lower level expression was detected in some

normal tissues when the cycle number was increased to 35. Antigen LST-S3-13 (SEQ ID NO: 66) was found to be expressed in 3 out of 4 lung tumors, one breast tumor and both colon tumor samples. Its expression in normal tissues was lower compared to tumors, and was only detected in 1 out of 4 normal lung tissues and in normal tissues from kidney, ovary and retina. Expression of antigens LST-S3-4 (SEQ ID NO: 62) and LST-S3-14 (SEQ ID NO: 67) was rare and did not show any tissue or tumor-specificity. Consistent with Northern blot analyses, the RT-PCT results on antigen LAT-S1-A-10A (SEQ ID NO: 78) suggested that its expression is high in lung, colon, stomach and small intestine tissues, including lung and colon tumors, whereas its expression was low or undetectable in other tissues.

A total of 2002 cDNA fragments isolated in lung subtractions I, II and III, described above, were colony PCR amplified and their mRNA expression levels in lung tumor, normal lung, and various other normal and tumor tissues were determined using microarray technology (Synteni, Palo Alto, CA). Briefly, the PCR amplification products were dotted onto slides in an array format, with each product occupying a unique location in the array. mRNA was extracted from the tissue sample to be tested, reverse transcribed, and fluorescent-labeled cDNA probes were generated. The microarrays were probed with the labeled cDNA probes, the slides scanned and fluorescence intensity was measured. This intensity correlates with the hybridization intensity. Seventeen non-redundant cDNA clones showed over-expression in lung squamous tumors, with expression in normal tissues tested (lung, skin, lymph node, colon, liver, pancreas, breast, heart, bone marrow, large intestine, kidney, stomach, brain, small intestine, bladder and salivary gland) being either undetectable, or 10-fold less compared to lung squamous tumors. The determined partial cDNA sequences for the clone L513S are provided in SEQ ID NO: 87 and 88; those for L514S are provided in SEQ ID NO: 89 and 90; those for L516S in SEQ ID NO: 91 and 92; that for L517S in SEQ ID NO: 93; that for L519S in SEQ ID NO: 94; those for L520S in SEQ ID NO: 95 and 96; those for L521S in SEQ ID NO: 97 and 98; that for L522S in SEQ ID NO: 99; that for L523S in SEQ ID NO: 100; that for L524S in SEQ ID NO: 101; that for L525S in SEQ ID NO: 102; that for L526S in SEQ ID NO: 103; that for L527S in SEQ ID NO: 104; that for L528S in SEQ ID NO: 105; that for L529S in SEQ ID NO: 106;

and those for L530S in SEQ ID NO: 107 and 108. Additionally, the full-length cDNA sequence for L530S is provided in SEQ ID NO: 151, with the corresponding predicted amino acid sequence being provided in SEQ ID NO: 152. L530S shows homology to a splice variant of a p53 tumor suppressor homologue, p63. The cDNA sequences of 7 known isoforms of p63 are provided in SEQ ID NO: 331-337, with the corresponding predicted amino acid sequences being provided in SEQ ID NO: 338-344, respectively.

Due to polymorphisms, the clone L531S appears to have two forms. A first determined full-length cDNA sequence for L531S is provided in SEQ ID NO: 109, with the corresponding predicted amino acid sequence being provided in SEQ ID NO: 110. A second determined full-length cDNA sequence for L531S is provided in SEQ ID NO: 111, with the corresponding predicted amino acid sequence being provided in SEQ ID NO: 112. The sequence of SEQ ID NO: 111 is identical to that of SEQ ID NO: 109, except that it contains a 27 bp insertion. Similarly, L514S also has two alternatively spliced forms; the first variant cDNA is listed as SEQ ID NO: 153, with the corresponding amino acid sequence being provided in SEQ ID NO: 155. The second variant form of L514S full-length cDNA is provided in SEQ ID NO: 154, with its corresponding amino acid sequence being provided in SEQ ID NO: 156.

Full length cloning for L524S (SEQ ID NO: 101) yielded two variants (SEQ ID NO: 163 and 164) with the corresponding predicted amino acid sequences of SEQ ID NO: 165 and 166, respectively. Both variants have been shown to encode parathyroid hormone-related peptide.

Attempts to isolate the full-length cDNA for L519S, resulted in the isolation of the extended cDNA sequence provided in SEQ ID NO: 173, which contains a potential open reading frame. The predicted amino acid sequence encoded by the sequence of SEQ ID NO: 173 is provided in SEQ ID NO: 174. Additionally, the full-length cDNA sequence for the clone of SEQ ID NO: 100 (known as L523S), a known gene, is provided in SEQ ID NO: 175, with the corresponding predicted amino acid sequence provided in SEQ ID NO: 176. In further studies, a full-length cDNA sequence for L523S was isolated from a L523S-positive tumor cDNA library by PCR amplification using gene specific primers designed from the sequence of SEQ ID NO: 175. The determined cDNA sequence is provided in SEQ ID NO: **. The amino acid

sequence encoded by this sequence is provided in SEQ ID NO: **. This protein sequence differs from the previously published protein sequence at two amino acid positions, namely at positions 158 and 410.

Comparison of the sequences of L514S and L531S (SEQ ID NO: 87 and 88, 89 and 90, and 109, respectively) with those in the gene bank, as described above, revealed no significant homologies to known sequences. The sequences of L513S, L516S, L517S, L519S, L520S and L530S (SEQ ID NO: 87 and 88, 91 and 92, 93, 94, 95 and 96, 107 and 108, respectively) were found to show some homology to previously identified ESTs. The sequences of L521S, L522S, L523S, L524S, L525S, L526S, L527S, L528S and L529S (SEQ ID NO: 97 and 98, 99, 99, 101, 102, 103, 104, 105, and 106, respectively) were found to represent known genes. The determined full-length cDNA sequences for L520S is provided in SEQ ID NO: 113, with the corresponding predicted amino acid sequence being provided in SEQ ID NO: 114. Subsequent microarray analysis has shown L520S to be overexpressed in breast tumors in addition to lung squamous tumors.

Further analysis has demonstrated that L529S (SEQ ID NO: 106 and 115), L525S (SEQ ID NO: 102 and 120) and L527S (SEQ ID NO: 104) are cytoskeletal components and potentially squamous cell specific proteins. L529S is connexin 26, a gap junction protein. It is highly expressed in lung squamous tumor 9688T, and moderately over-expressed in two others. However, lower level expression of connexin 26 is also detectable in normal skin, colon, liver and stomach. The over-expression of connexin 26 in some breast tumors has been reported and a mutated form of L529S may result in over-expression in lung tumors. L525S is plakophilin 1, a desmosomal protein found in plaque-bearing adhering junctions of the skin. Expression levels for L525S mRNA is highly elevated in three out of four lung squamous tumors tested, and in normal skin. L527S has been identified as keratin 6 isoform, type II 58 Kd keratin, and cytokeratin 13 and shows over-expression in squamous tumors and low expression in normal skin, breast and colon tissues. Notably, keratin and keratin-related genes have been extensively documented as potential markers for lung cancer including CYFRA2.1 (Pastor, A., et al, *Eur. Respir. J.*, 10:603-609, 1997). L513S (SEQ ID NO: 87 and 88)

shows moderate over-expression in several tumor tissues tested, and encodes a protein that was first isolated as a pemphigus vulgaris antigen.

L520S (SEQ ID NO: 95 and 96) and L521S (SEQ ID NO: 97 and 98) are highly expressed in lung squamous tumors, and L520S is up-regulated in normal salivary gland and L521S is over-expressed in normal skin. Both belong to a family of small proline rich proteins and represent markers for fully differentiated squamous cells. L521S has been described as a specific marker for lung squamous tumor (Hu, R., et al, *Lung Cancer*, 20:25-30, 1998). L515S (SEQ ID NO: 162) encodes IGF- β 2 and L516S is an aldose reductase homologue and both are moderately expressed in lung squamous tumors and in normal colon. Notably, L516S (SEQ ID NO: 91 and 92) is up-regulated in metastatic tumors but not primary lung adenocarcinoma, an indication of its potential role in metatasis and a potential prognostic marker. L522S (SEQ ID NO: 99) is moderately over-expressed in lung squamous tumors with minimum expression in normal tissues. L522S has been shown to belong to a class IV alcohol dehydrogenase, ADH7, and its expression profile suggests it is a squamous cell specific antigen. L523S (SEQ ID NO: 100) is moderately over-expressed in lung squamous tumor, human pancreatic cancer cell lines and pancreatic cancer tissues, suggesting this gene may be a shared antigen between pancreatic and lung squamous cell cancer.

L524S (SEQ ID NO: 101) is over-expressed in the majority of squamous tumors tested and is homologous with parathyroid hormone-related peptide (PTHrP), which is best known to cause humoral hypercalcaemia associated with malignant tumors such as leukemia, prostate and breast cancer. It is also believed that PTHrP is most commonly associated with squamous carcinoma of lung and rarely with lung adenocarcinoma (Davidson, L.A., et al, *J. Pathol.*, 178: 398-401, 1996). L528S (SEQ ID NO: 105) is highly over-expressed in two lung squamous tumors with moderate expression in two other squamous tumors, one lung adenocarcinoma and some normal tissues, including skin, lymph nodes, heart, stomach and lung. It encodes the NMB gene that is similar to the precursor of melanocyte specific gene Pmel17, which is reported to be preferentially expressed in low-metastatic potential melanoma cell lines. This suggests that L528S may be a shared antigen in both melanoma and lung squamous cell carcinoma. L526S (SEQ ID NO: 103) is overexpressed in all lung

squamous cell tumor tissues tested and has been shown to share homology with a gene (ATM) in which a mutation causes ataxia telangiectasia, a genetic disorder in humans causing a predisposition to cancer, among other symptoms. ATM encodes a protein that activates p53 mediated cell-cycle checkpoint through direct binding and phosphorylation of the p53 molecule. Approximately 40% of lung cancer is associated with p53 mutations, and it is speculated that over-expression of ATM is a result of compensation for loss of p53 function, but it is unknown whether over-expression is the cause of result of lung squamous cell carcinoma. Additionally, expression of L526S (ATM) is also detected in a metastatic but not lung adenocarcinoma, suggesting a role in metastasis.

Expression of L523S (SEQ ID NO: 175), was also examined by real time RT-PCR as described above. In a first study using a panel of lung squamous tumors, L523S was found to be expressed in 4/7 lung squamous tumors, 2/3 head and neck squamous tumors and 2/2 lung adenocarcinomas, with low level expression being observed in skeletal muscle, soft palate and tonsil. In a second study using a lung adenocarcinoma panel, expression of L523S was observed in 4/9 primary adenocarcinomas, 2/2 lung pleural effusions, 1/1 metastatic lung adenocarcinomas and 2/2 lung squamous tumors, with little expression being observed in normal tissues.

Expression of L523S in lung tumors and various normal tissues was also examined by Northern blot analysis, using standard techniques. In a first study, L523S was found to be expressed in a number of lung adenocarcinomas and squamous cell carcinomas, as well as normal tonsil. No expression was observed in normal lung. In a second study using a normal tissue blot (HB-12) from Clontech, no expression was observed in brain, skeletal muscle, colon, thymus, spleen, kidney, liver, small intestine, lung or PBMC, although there was strong expression in placenta.

EXAMPLE 3

ISOLATION AND CHARACTERIZATION OF LUNG TUMOR POLYPEPTIDES BY PCR-BASED SUBTRACTION

Eight hundred and fifty seven clones from a cDNA subtraction library, containing cDNA from a pool of two human lung squamous tumors subtracted against eight normal human tissue cDNAs including lung, PBMC, brain, heart, kidney, liver, pancreas, and skin, (Clontech, Palo Alto, CA) were derived and submitted to a first round of PCR amplification. This library was subjected to a second round of PCR amplification, following the manufacturer's protocol. The resulting cDNA fragments were subcloned into the vector P7- Adv vector (Clontech, Palo Alto, CA) and transformed into DH5 α *E. coli* (Gibco, BRL). DNA was isolated from independent clones and sequenced using a Perkin Elmer/Applied Biosystems Division Automated Sequencer Model 373A.

One hundred and sixty two positive clones were sequenced. Comparison of the DNA sequences of these clones with those in the the EMBL and GenBank databases, as described above, revealed no significant homologies to 13 of these clones, hereinafter referred to as Contigs 13, 16, 17, 19, 22, 24, 29, 47, 49, 56-59. The determined cDNA sequences for these clones are provided in SEQ ID NO: 125, 127-129, 131-133, 142, 144, 148-150, and 157, respectively. Contigs 1, 3-5, 7-10, 12, 11, 15, 20, 31, 33, 38, 39, 41, 43, 44, 45, 48, 50, 53, 54 (SEQ ID NO: 115-124, 126, 130, 134-141, 143, 145-147, respectively) were found to show some degree of homology to previously identified DNA sequences. Contig 57 (SEQ ID NO: 149) was found to represent the clone L519S (SEQ ID NO: 94) disclosed in US. Patent Application No. 09/123,912, filed July 27, 1998. To the best of the inventors' knowledge, none of these sequences have been previously shown to be differentially over-expressed in lung tumors.

mRNA expression levels for representative clones in lung tumor tissues, normal lung tissues (n=4), resting PBMC, salivary gland, heart, stomach, lymph nodes, skeletal muscle, soft palate, small intestine, large intestine, bronchial, bladder, tonsil, kidney, esophagus, bone marrow, colon, adrenal gland, pancreas, and skin, (all derived from human) were determined by RT-PCR as described above. Expression levels using microarray technology, as described above, were examined in one sample of each tissue type unless otherwise indicated.

Contig 3 (SEQ ID NO: 116) was found to be highly expressed in all head and neck squamous cell tumors tested (17/17), and expressed in the majority (8/12) of lung squamous tumors, (high expression in 7/12, moderate in 2/12, and low in 2/12), while showing negative expression for 2/4 normal lung tissues and low expression in the remaining two samples. Contig 3 showed moderate expression in skin and soft palate, and lowered expression levels in resting PBMC, large intestine, salivary gland, tonsil, pancreas, esophagus, and colon. Contig 11 (SEQ ID NO: 124) was found to be expressed in all head and neck squamous cell tumors tested (17/17): highly expressed in 14/17, and moderately expressed in 3/17. Additionally, expression in lung squamous tumors showed high expression in 3/12 and moderate in 4/12. Contig 11 was negative for 3/4 normal lung samples, with the remaining sample having only low expression. Contig 11 showed low to moderate reactivity to salivary gland, soft palate, bladder, tonsil, skin, esophagus, and large intestine. Contig 13 (SEQ ID NO: 125) was found to be expressed in all head and neck squamous cell tumors tested (17/17): highly expressed in 12/17, and moderately expressed in 5/17. Contig 13 was expressed in 7/12 lung squamous tumors, with high expression in 4/12 and moderate expression in three samples. Analysis of normal lung samples showed negative expression for 2/4 and low to moderate expression in the remaining two samples. Contig 13 did show low to moderate reactivity to resting PBMC, salivary gland, bladder, pancreas, tonsil, skin, esophagus, and large intestine, as well as high expression in soft palate. Contig 16 (SEQ ID NO: 127) was found to be moderately expressed in some head and neck squamous cell tumors (6/17) and one lung squamous tumor; while showing no expression in any normal lung samples tested. Contig 16 did show low reactivity to resting PBMC, large intestine, skin, salivary gland, and soft palate. Contig 17 (SEQ ID NO: 128) was shown to be expressed in all head and neck squamous cell tumors tested (17/17): highly expressed in 5/17, and moderately expressed in 12/17. Expression levels in lung squamous tumors showed one tumor sample with high expression and 3/12 with moderate levels. Contig 17 was negative for 2/4 normal lung samples, with the remaining samples having only low expression. Additionally, low level expression was found in esophagus and soft palate. Contig 19 (SEQ ID NO: 129) was found to be expressed in most head and neck squamous cell tumors tested (11/17); with two

samples having high levels, 6/17 showing moderate expression, and low expression being found in 3/17. Testing in lung squamous tumors revealed only moderate expression in 3/12 samples. Expression levels in 2/4 of normal lung samples were negative, the two other samples having only low expression. Contig 19 showed low expression levels in esophagus, resting PBMC, salivary gland, bladder, soft palate and pancreas.

Contig 22 (SEQ ID NO: 131), was shown to be expressed in most head and neck squamous cell tumors tested (13/17) with high expression in four of these samples, moderate expression in 6/17, and low expression in 3/17. Expression levels in lung squamous tumors were found to be moderate to high for 3/12 tissues tested, with negative expression in two normal lung samples and low expression in two other samples (n=4). Contig 22 showed low expression in skin, salivary gland and soft palate. Similarly, Contig 24 (SEQ ID NO: 132) was found to be expressed in most head and neck squamous cell tumors tested (13/17) with high expression in three of these samples, moderate expression in 6/17, and low expression in 4/17. Expression levels in lung squamous tumors were found to be moderate to high for 3/12 tissues tested, with negative expression for three normal lung samples and low expression in one sample (n=4). Contig 24 showed low expression in skin, salivary gland and soft palate. Contig 29 (SEQ ID NO: 133) was expressed in nearly all head and neck squamous cell tumors tested (16/17): highly expressed in 4/17, moderately expressed in 11/17, with low expression in one sample. Also, it was moderately expressed in 3/12 lung squamous tumors, while being negative for 2/4 normal lung samples. Contig 29 showed low to moderate expression in large intestine, skin, salivary gland, pancreas, tonsil, heart and soft palate. Contig 47 (SEQ ID NO: 142) was expressed in most head and neck squamous cell tumors tested (12/17): moderate expression in 10/17, and low expression in two samples. In lung squamous tumors, it was highly expressed in one sample and moderately expressed in two others (n=13). Contig 47 was negative for 2/4 normal lung samples, with the remaining two samples having moderate expression. Also, Contig 47 showed moderate expression in large intestine, and pancreas, and low expression in skin, salivary gland, soft palate, stomach, bladder, resting PBMC, and tonsil.

Contig 48 (SEQ ID NO: 143) was expressed in all head and neck squamous cell tumors tested (17/17): highly expressed in 8/17 and moderately expressed in 7/17, with low expression in two samples. Expression levels in lung squamous tumors were high to moderate in three samples (n=13). Contig 48 was negative for one out of four normal lung samples, the remaining showing low or moderate expression. Contig 48 showed moderate expression in soft palate, large intestine, pancreas, and bladder, and low expression in esophagus, salivary gland, resting PBMC, and heart. Contig 49 (SEQ ID NO: 144) was expressed at low to moderate levels in 6/17 head and neck squamous cell tumors tested. Expression levels in lung squamous tumors were moderate in three samples (n=13). Contig 49 was negative for 2/4 normal lung samples, the remaining samples showing low expression. Moderate expression levels in skin, salivary gland, large intestine, pancreas, bladder and resting PBMC were shown, as well as low expression in soft palate, lymph nodes, and tonsil. Contig 56 (SEQ ID NO: 148) was expressed in low to moderate levels in 3/17 head and neck squamous cell tumors tested, and in lung squamous tumors, showing low to moderate levels in three out of thirteen samples. Notably, low expression levels were detected in one adenocarcinoma lung tumor sample (n=2). Contig 56 was negative for 3/4 normal lung samples, and showed moderate expression levels in only large intestine, and low expression in salivary gland, soft palate, pancreas, bladder, and resting PBMC. Contig 58, also known as L769P, (SEQ ID NO: 150) was expressed at moderate levels in 11/17 head and neck squamous cell tumors tested and low expression in one additional sample. Expression in lung squamous tumors showed low to moderate levels in three out of thirteen samples. Contig 58 was negative for 3/4 normal lung samples, with one sample having low expression. Moderate expression levels in skin, large intestine, and resting PBMC were demonstrated, as well as low expression in salivary gland, soft palate, pancreas, and bladder. Contig 59 (SEQ ID NO: 157) was expressed in some head, neck, and lung squamous tumors. Low level expression of Contig 59 was also detected in salivary gland and large intestine.

The full-length cDNA sequence for Contig 22, also referred to as L763P, is provided in SEQ ID NO: 158, with the corresponding predicted amino acid sequence being provided in SEQ ID NO: 159. Real-time RT-PCR analysis of L763P revealed

that it is highly expressed in 3/4 lung squamous tumors as well as 4/4 head and neck squamous tumors, with low level expression being observed in normal brain, skin, soft pallet and trachea. Subsequent database searches revealed that the sequence of SEQ ID NO: 158 contains a mutation, resulting in a frameshift in the corresponding protein sequence. A second cDNA sequence for L763P is provided in SEQ ID NO: 345, with the corresponding amino acid sequence being provided in SEQ ID NO: 346. The sequences of SEQ ID NO: 159 and 346 are identical with the exception of the C-terminal 33 amino acids of SEQ ID NO: 159.

The full-length cDNA sequence incorporating Contigs 17, 19, and 24, referred to as L762P, is provided in SEQ ID NO: 160, with the corresponding predicted amino acid sequence being provided in SEQ ID NO: 161. Further analysis of L762P has determined it to be a type I membrane protein and two additional variants have been sequenced. Variant 1 (SEQ ID NO: 167, with the corresponding amino acid sequence in SEQ ID NO: 169) is an alternatively spliced form of SEQ ID NO: 160 resulting in deletion of 503 nucleotides, as well as deletion of a short segment of the expressed protein. Variant 2 (SEQ ID NO: 168, with the corresponding amino acid sequence in SEQ ID NO: 170) has a two nucleotide deletion at the 3' coding region in comparison to SEQ ID NO: 160, resulting in a secreted form of the expressed protein. Real-time RT-PCR analysis of L762P revealed that is over-expressed in 3/4 lung squamous tumors and 4/4 head & neck tumors, with low level expression being observed in normal skin, soft pallet and trachea.

The full-length cDNA sequence for contig 56 (SEQ ID NO: 148), also referred to as L773P, is provided in SEQ ID NO: 171, with the predicted amino acid sequence in SEQ ID NO: 172. L773P was found to be identical to dihydroxyl dehydrogenase at the 3' portion of the gene, with divergent 5' sequence. As a result, the 69 N-terminal amino acids are unique. The cDNA sequence encoding the 69 N-terminal amino acids is provided in SEQ ID NO: 349, with the N-terminal amino acid sequence being provided in SEQ ID NO: 350. Real-time PCR revealed that L773P is highly expressed in lung squamous tumor and lung adenocarcinoma, with no detectable expression in normal tissues. Subsequent Northern blot analysis of L773P demonstrated that this transcript is differentially over-expressed in squamous tumors

and detected at approximately 1.6 Kb in primary lung tumor tissue and approximately 1.3 Kb in primary head and neck tumor tissue.

Subsequent microarray analysis has shown Contig 58, also referred to as L769S (SEQ ID NO: 150), to be overexpressed in breast tumors in addition to lung squamous tumors.

EXAMPLE 4

SYNTHESIS OF POLYPEPTIDES

Polypeptides may be synthesized on a Perkin Elmer/Applied Biosystems Division 430A peptide synthesizer using Fmoc chemistry with HPTU (O-Benzotriazole-N,N,N',N'-tetramethyluronium hexafluorophosphate) activation. A Gly-Cys-Gly sequence may be attached to the amino terminus of the peptide to provide a method of conjugation, binding to an immobilized surface, or labeling of the peptide. Cleavage of the peptides from the solid support may be carried out using the following cleavage mixture: trifluoroacetic acid:ethanedithiol:thioanisole:water:phenol (40:1:2:2:3). After cleaving for 2 hours, the peptides may be precipitated in cold methyl-t-butyl-ether. The peptide pellets may then be dissolved in water containing 0.1% trifluoroacetic acid (TFA) and lyophilized prior to purification by C18 reverse phase HPLC. A gradient of 0%-60% acetonitrile (containing 0.1% TFA) in water (containing 0.1% TFA) may be used to elute the peptides. Following lyophilization of the pure fractions, the peptides may be characterized using electrospray or other types of mass spectrometry and by amino acid analysis.

EXAMPLE 5

PREPARATION OF ANTIBODIES AGAINST LUNG CANCER ANTIGENS

Polyclonal antibodies against the lung cancer antigens L514S, L528S and L531S (SEQ ID NO: 155, 225 and 112, respectively) were prepared as follows.

Rabbits were immunized with recombinant protein expressed in and purified from *E. coli* as described above. For the initial immunization, 400 µg of

antigen combined with muramyl dipeptide (MDP) was injected subcutaneously (S.C.). Animals were boosted S.C. 4 weeks later with 200 µg of antigen mixed with incomplete Freund's Adjuvant (IFA). Subsequent boosts of 100 µg of antigen mixed with IFA were injected S.C. as necessary to induce high antibody titer responses. Serum bleeds
5 from immunized rabbits were tested for antigen-specific reactivity using ELISA assays with purified protein. Polyclonal antibodies against L514S, L528S and L531S were affinity purified from high titer polyclonal sera using purified protein attached to a solid support.

Immunohistochemical analysis using polyclonal antibodies against
10 L514S was performed on a panel of 5 lung tumor samples, 5 normal lung tissue samples and normal colon, kidney, liver, brain and bone marrow. Specifically, tissue samples were fixed in formalin solution for 24 hours and embedded in paraffin before being sliced into 10 micron sections. Tissue sections were permeabilized and incubated with antibody for 1 hr. HRP-labeled anti-mouse followed by incubation with DAB
15 chromogen was used to visualize L514S immunoreactivity. L514S was found to be highly expressed in lung tumor tissue with little or no expression being observed in normal lung, brain or bone marrow. Light staining was observed in colon and kidney. Staining was seen in normal liver but no mRNA has been detected in this tissue making this result suspect.

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EXAMPLE 6

PEPTIDE PRIMING OF MICE AND PROPAGATION OF CTL LINES

Immunogenic peptides from the lung cancer antigen L762P (SEQ ID
25 NO: 161) for HLA-A2/K^b-restricted CD8⁺ T cells were identified as follows.

The location of HLA-A2 binding peptides within the lung cancer antigen L762P (SEQ ID NO: 161) was predicted using a computer program which predicts peptides sequences likely to be to HLA-A*0201 by fitting to the known peptide binding motif for HLA-A*0201 (Rupert *et al.* (1993) *Cell* 74:929; Rammensee *et al.*
30 (1995) *Immunogenetics* 41:178-228). A series of 19 synthetic peptides corresponding to a selected subset of the predicted HLA-A*0201 binding peptides was prepared as described above.

Mice expressing the transgene for human HLA A2/K^b (provided by Dr L. Sherman, The Scripps Research Institute, La Jolla, CA) were immunized with the synthetic peptides, as described by Theobald et al., *Proc. Natl. Acad. Sci. USA* 92:11993-11997, 1995 with the following modifications. Mice were immunized with 50µg of L726P peptide and 120µg of an I-A^b binding peptide derived from hepatitis B Virus protein emulsified in incomplete Freund's adjuvant. Three weeks later these mice were sacrificed and single cell suspensions prepared. Cells were then resuspended at 7×10^6 cells/ml in complete media (RPMI-1640; Gibco BRL, Gaithersburg, MD) containing 10% FCS, 2mM Glutamine (Gibco BRL), sodium pyruvate (Gibco BRL), non-essential amino acids (Gibco BRL), 2×10^{-5} M 2-mercaptoethanol, 50U/ml penicillin and streptomycin, and cultured in the presence of irradiated (3000 rads) L762P peptide- (5µg/ml) and 10mg/ml B₂-microglobulin- (3 µg/ml) LPS blasts (A2 transgenic spleens cells cultured in the presence of 7µg/ml dextran sulfate and 25µg/ml LPS for 3 days). After six days, cells (5×10^5 /ml) were restimulated with 2.5×10^6 /ml peptide pulsed irradiated (20,000 rads) EL4A2Kb cells (Sherman et al, *Science* 258:815-818, 1992) and 5×10^6 /ml irradiated (3000 rads) A2/K^b-transgenic spleen feeder cells. Cells were cultured in the presence of 10U/ml IL-2. Cells were restimulated on a weekly basis as described, in preparation for cloning the line.

Peptide-specific cell lines were cloned by limiting dilution analysis with irradiated (20,000 rads) L762P peptide-pulsed EL4 A2Kb tumor cells (1×10^4 cells/well) as stimulators and irradiated (3000 rads) A2/K^b-transgenic spleen cells as feeders (5×10^5 cells/ well) grown in the presence of 10U/ml IL-2. On day 7, cells were restimulated as before. On day 14, clones that were growing were isolated and maintained in culture.

Cell lines specific for L762P-87 (SEQ ID NO: 226; corresponding to amino acids 87-95 of SEQ ID NO: 161), L726P-145 (SEQ ID NO: 227; corresponding to amino acids 145-153 of SEQ ID NO: 161), L726P-585 (SEQ ID NO: 228; corresponding to amino acids 585-593 of SEQ ID NO: 161), L762P-425 (SEQ ID NO: 229; corresponding to amino acids 425-433 of SEQ ID NO: 161), L762P(10)-424 (SEQ ID NO: 230; corresponding to amino acids 424-433 of SEQ ID NO: 161) and L762P(10)-458 (SEQ ID NO: 231; corresponding to amino acids 458-467 of SEQ ID

NO: 161) demonstrated significantly higher reactivity (as measured by percent specific lysis) against L762P peptide-pulsed EL4-A2/K^b tumor target cells than control peptide-pulsed EL4-A2/K^b tumor target cells.

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EXAMPLE 7

IDENTIFICATION OF CD4 IMMUNOGENIC T CELL EPITOPES DERIVED
FROM THE LUNG CANCER ANTIGEN L762P

CD4 T cell lines specific for the antigen L762P (SEQ ID NO: 161) were
10 generated as follows.

A series of 28 overlapping peptides were synthesized that spanned approximately 50% of the L762P sequence. For priming, peptides were combined into pools of 4-5 peptides, pulsed at 20 micrograms/ml into dendritic cells for 24 hours. The dendritic cells were then washed and mixed with positively selected CD4⁺ T cells in 96
15 well U-bottomed plates. Forty cultures were generated for each peptide pool. Cultures were restimulated weekly with fresh dendritic cells loaded with peptide pools. Following a total of 3 stimulation cycles, cells were rested for an additional week and tested for specificity to antigen presenting cells (APC) pulsed with peptide pools using interferon-gamma ELISA and proliferation assays. For these assays, adherent
20 monocytes loaded with either the relevant peptide pool or an irrelevant peptide were used as APC. T cell lines that appeared to specifically recognize L762P peptide pools both by cytokine release and proliferation were identified for each pool. Emphasis was placed on identifying T cells with proliferative responses. T cell lines that demonstrated either both L762P-specific cytokine secretion and proliferation, or strong proliferation
25 alone were further expanded to be tested for recognition of individual peptides from the pools, as well as for recognition of recombinant L762P. The source of recombinant L762P was *E. coli*, and the material was partially purified and endotoxin positive. These studies employed 10 micrograms of individual peptides, 10 or 2 micrograms of an irrelevant peptide, and 2 or 0.5 micrograms of either L762P protein or an irrelevant,
30 equally impure, *E. coli* generated recombinant protein. Significant interferon-gamma production and CD4 T cell proliferation was induced by a number of L762P-derived

peptides in each pool. The amino acid sequences for these peptides are provided in SEQ ID NO: 232-251. These peptides correspond to amino acids 661-680, 676-696, 526-545, 874-893, 811-830, 871-891, 856-875, 826-845, 795-815, 736-755, 706-725, 706-725, 691-710, 601-620, 571-590, 556-575, 616-635, 646-665, 631-650, 541-560 and 586-605, respectively, of SEQ ID NO: 161.

CD4 T cell lines that demonstrated specificity for individual L762P-derived peptides were further expanded by stimulation with the relevant peptide at 10 micrograms/ml. Two weeks post-stimulation, T cell lines were tested using both proliferation and IFN-gamma ELISA assays for recognition of the specific peptide. A number of previously identified T cells continued to demonstrate L762P-peptide specific activity. Each of these lines was further expanded on the relevant peptide and, following two weeks of expansion, tested for specific recognition of the L762P-peptide in titration experiments, as well as for recognition of recombinant *E. coli*-derived L762P protein. For these experiments, autologous adherent monocytes were pulsed with either the relevant L762P-derived peptide, an irrelevant mammaglobin-derived peptide, recombinant *E. coli*-derived L762P (approx. 50% pure), or an irrelevant *E. coli*-derived protein. The majority of T cell lines were found to show low affinity for the relevant peptide, since specific proliferation and IFN-gamma ratios dramatically decreased as L762P peptide was diluted. However, four lines were identified that demonstrated significant activity even at 0.1 micrograms/ml peptide. Each of these lines (referred to as A/D5, D/F5, E/A7 and E/B6) also appeared to specifically proliferate in response to the *E. coli*-derived L762P protein preparation, but not in response to the irrelevant protein preparation. The amino acid sequences of the L762P-derived peptides recognized by these lines are provided in SEQ ID NO: 234, 249, 236 and 245, respectively. No protein specific IFN-gamma was detected for any of the lines. Lines A/D5, E/A7 and E/B6 were cloned on autologous adherent monocytes pulsed with the relevant peptide at 0.1 (A/D5 and E/A7) or 1 (D/F5) microgram/ml. Following growth, clones were tested for specificity for the relevant peptide. Numerous clones specific for the relevant peptide were identified for lines A/D5 and E/A7.

EXAMPLE 8**PROTEIN EXPRESSION OF LUNG TUMOR-SPECIFIC ANTIGENS****5 a) Expression of L514S in *E. coli***

The lung tumor antigen L514S (SEQ ID NO: 89) was subcloned into the expression vector pE32b at NcoI and NotI sites, and transformed into *E. coli* using standard techniques. The protein was expressed from residues 3-153 of SEQ ID NO: 89. The expressed amino acid sequence and the corresponding DNA sequence are
10 provided in SEQ ID NO: 252 and 253, respectively.

b) Expression of L762P

Amino acids 32-944 of the lung tumor antigen L762P (SEQ ID NO: 161), with a 6X His Tag, were subcloned into a modified pET28 expression vector,
15 using kanamycin resistance, and transformed into BL21 CodonPlus using standard techniques. Low to moderate levels of expression were observed. The determined DNA sequence of the L762P expression construct is provided in SEQ ID NO: 254.

From the foregoing it will be appreciated that, although specific
20 embodiments of the invention have been described herein for purposes of illustration, various modifications may be made without deviating from the spirit and scope of the invention. Accordingly, the invention is not limited except as by the appended claims.

CLAIMS

1. An isolated polypeptide, comprising at least an immunogenic portion of a lung tumor protein, or a variant thereof, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide sequence selected from the group consisting of:

(a) sequences recited in SEQ ID NO: 1-3, 6-8, 10-13, 15-27, 29, 30, 32, 34-49, 51, 52, 54, 55, 57-59, 61-69, 71, 73, 74, 77, 78, 80-82, 84, 86-96, 107-109, 111, 113, 125, 127, 128, 129, 131-133, 142, 144, 148-151, 153, 154, 157, 158, 160, 167, 168, 171, 173, 175, 179, 182, 184-186, 188-191, 193, 194, 198-207, 209, 210, 213, 214, 217, 220-224, 253, 254-258, 260, 262-264, 270, 272, 275, 276, 279-281, 286, 287, 291, 293, 295, 296, 300, 302, 308-310, 313, 315-317, 323, 345, 347 and 349;

(b) sequences that hybridize to a sequence recited in any one of SEQ ID NO: 1-3, 6-8, 10-13, 15-27, 29, 30, 32, 34-49, 51, 52, 54, 55, 57-59, 61-69, 71, 73, 74, 77, 78, 80-82, 84, 86-96, 107-109, 111, 113, 125, 127, 128, 129, 131-133, 142, 144, 148-151, 153, 154, 157, 158, 160, 167, 168, 171, 173, 175, 179, 182, 184-186, 188-191, 193, 194, 198-207, 209, 210, 213, 214, 217, 220-224, 253, 254-258, 260, 262-264, 270, 272, 275, 276, 279-281, 286, 287, 291, 293, 295, 296, 300, 302, 308-310, 313, 315-317, 323, 345, 347 and 349 under moderately stringent conditions; and

(c) complements of sequences of (a) or (b).

2. An isolated polypeptide according to claim 1, wherein the polypeptide comprises an amino acid sequence that is encoded by a polynucleotide sequence recited in any one of SEQ ID NO: 1-3, 6-8, 10-13, 15-27, 29, 30, 32, 34-49, 51, 52, 54, 55, 57-59, 61-69, 71, 73, 74, 77, 78, 80-82, 84, 86-96, 107-109, 111, 113, 125, 127, 128, 129, 131-133, 142, 144, 148-151, 153, 154, 157, 158,

160, 167, 168, 171, 173, 175, 179, 182, 184-186, 188-191, 193, 194, 198-207, 209, 210, 213, 214, 217, 220-224, 253, 254-258, 260, 262-264, 270, 272, 275, 276, 279-281, 286, 287, 291, 293, 295, 296, 300, 302, 308-310, 313, 315-317, 323, 345, 347 and 349 or a complement of any of the foregoing polynucleotide sequences.

5

3. An isolated polypeptide comprising a sequence recited in any one of SEQ ID NO: 110, 112, 114, 152, 155, 156, 159, 161, 165, 166, 169, 170, 172, 174, 176, 226-252, 346, 348 and 350.

4. An isolated polynucleotide encoding at least 15 amino acid
10 residues of a lung tumor protein, or a variant thereof that differs in one or more substitutions, deletions, additions and/or insertions such that the ability of the variant to react with antigen-specific antisera is not substantially diminished, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide comprising a sequence recited in any one of SEQ ID NO: 1-3, 6-8, 10-13, 15-27, 29,
15 30, 32, 34-49, 51, 52, 54, 55, 57-59, 61-69, 71, 73, 74, 77, 78, 80-82, 84, 86-96, 107-109, 111, 113, 125, 127, 128, 129, 131-133, 142, 144, 148-151, 153, 154, 157, 158, 160, 167, 168, 171, 173, 175, 179, 182, 184-186, 188-191, 193, 194, 198-207, 209, 210, 213, 214, 217, 220-224, 253, 254-258, 260, 262-264, 270, 272, 275, 276, 279-281, 286, 287, 291, 293, 295, 296, 300, 302, 308-310, 313, 315-317, 323, 345, 347 and 349 or a
20 complement of any of the foregoing sequences.

5. An isolated polynucleotide encoding a lung tumor protein, or a variant thereof, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide comprising a sequence recited in any one of SEQ ID NO:
25 1-3, 6-8, 10-13, 15-27, 29, 30, 32, 34-49, 51, 52, 54, 55, 57-59, 61-69, 71, 73, 74, 77, 78, 80-82, 84, 86-96, 107-109, 111, 113, 125, 127, 128, 129, 131-133, 142, 144, 148-151, 153, 154, 157, 158, 160, 167, 168, 171, 173, 175, 179, 182, 184-186, 188-191, 193, 194, 198-207, 209, 210, 213, 214, 217, 220-224, 253, 254-258, 260, 262-264, 270, 272, 275, 276, 279-281, 286, 287, 291, 293, 295, 296, 300, 302, 308-310, 313, 315-317,
30 323, 345, 347 and 349 or a complement of any of the foregoing sequences.

6. An isolated polynucleotide, comprising a sequence recited in any one of SEQ ID NO: 1-3, 6-8, 10-13, 15-27, 29, 30, 32, 34-49, 51, 52, 54, 55, 57-59, 61-69, 71, 73, 74, 77, 78, 80-82, 84, 86-96, 107-109, 111, 113, 125, 127, 128, 129, 131-133, 142, 144, 148-151, 153, 154, 157, 158, 160, 167, 168, 171, 173, 175, 179, 182, 184-186, 188-191, 193, 194, 198-207, 209, 210, 213, 214, 217, 220-224, 253, 254-258, 260, 262-264, 270, 272, 275, 276, 279-281, 286, 287, 291, 293, 295, 296, 300, 302, 308-310, 313, 315-317, 323, 345, 347 and 349.

7. An isolated polynucleotide, comprising a sequence that hybridizes to a sequence recited in any one of SEQ ID NO: 1-3, 6-8, 10-13, 15-27, 29, 30, 32, 34-49, 51, 52, 54, 55, 57-59, 61-69, 71, 73, 74, 77, 78, 80-82, 84, 86-96, 107-109, 111, 113, 125, 127, 128, 129, 131-133, 142, 144, 148-151, 153, 154, 157, 158, 160, 167, 168, 171, 173, 175, 179, 182, 184-186, 188-191, 193, 194, 198-207, 209, 210, 213, 214, 217, 220-224, 253, 254-258, 260, 262-264, 270, 272, 275, 276, 279-281, 286, 287, 291, 293, 295, 296, 300, 302, 308-310, 313, 315-317, 323, 345, 347 and 349 under moderately stringent conditions.

8. An isolated polynucleotide complementary to a polynucleotide according to any one of claims 4-7.

9. An expression vector, comprising a polynucleotide according to any one of claims claim 4-8.

10. A host cell transformed or transfected with an expression vector according to claim 9.

11. An isolated antibody, or antigen-binding fragment thereof, that specifically binds to a lung tumor protein that comprises an amino acid sequence that is encoded by a polynucleotide sequence recited in any one of SEQ ID NO: 1-3, 6-8, 10-13, 15-27, 29, 30, 32, 34-49, 51, 52, 54, 55, 57-59, 61-69, 71, 73, 74, 77, 78, 80-82, 84,

86-96, 107-109, 111, 113, 125, 127, 128, 129, 131-133, 142, 144, 148-151, 153, 154, 157, 158, 160, 167, 168, 171, 173, 175, 179, 182, 184-186, 188-191, 193, 194, 198-207, 209, 210, 213, 214, 217, 220-224, 253, 254-258, 260, 262-264, 270, 272, 275, 276, 279-281, 286, 287, 291, 293, 295, 296, 300, 302, 308-310, 313, 315-317, 323, 345, 347 and 5 349 or a complement of any of the foregoing polynucleotide sequences.

12. A fusion protein, comprising at least one polypeptide according to claim 1.

10 13. A fusion protein according to claim 12, wherein the fusion protein comprises an expression enhancer that increases expression of the fusion protein in a host cell transfected with a polynucleotide encoding the fusion protein.

14. A fusion protein according to claim 12, wherein the fusion 15 protein comprises a T helper epitope that is not present within the polypeptide of claim 1.

15. A fusion protein according to claim 12, wherein the fusion protein comprises an affinity tag.

20

16. An isolated polynucleotide encoding a fusion protein according to claim 12.

17. A pharmaceutical composition, comprising a physiologically 25 acceptable carrier and at least one component selected from the group consisting of:

- (a) a polypeptide according to claim 1;
- (b) a polynucleotide according to claim 4;
- (c) an antibody according to claim 11;
- (d) a fusion protein according to claim 12; and
- 30 (e) a polynucleotide according to claim 16.

18. A vaccine comprising an immunostimulant and at least one component selected from the group consisting of:
- (a) a polypeptide according to claim 1;
 - (b) a polynucleotide according to claim 4;
 - (c) an antibody according to claim 11;
 - (d) a fusion protein according to claim 12; and
 - (e) a polynucleotide according to claim 16.
19. A vaccine according to claim 18, wherein the immunostimulant is an adjuvant.
20. A vaccine according to any claim 18, wherein the immunostimulant induces a predominantly Type I response.
21. A method for inhibiting the development of a cancer in a patient, comprising administering to a patient an effective amount of a pharmaceutical composition according to claim 17.
22. A method for inhibiting the development of a cancer in a patient, comprising administering to a patient an effective amount of a vaccine according to claim 18.
23. A pharmaceutical composition comprising an antigen-presenting cell that expresses a polypeptide according to claim 1, in combination with a pharmaceutically acceptable carrier or excipient.
24. A pharmaceutical composition according to claim 23, wherein the antigen presenting cell is a dendritic cell or a macrophage.

25. A vaccine comprising an antigen-presenting cell that expresses a polypeptide comprising at least an immunogenic portion of a lung tumor protein, or a variant thereof, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide sequence selected from the group consisting of:

5 (a) sequences recited in SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349;

(b) sequences that hybridize to a sequence recited in any one of SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171,
10 173, 175, 177-224, 255-337, 345, 347 and 349 under moderately stringent conditions;
and

(c) complements of sequences of (i) or (ii);

in combination with an immunostimulant.

15 26. A vaccine according to claim 25, wherein the immunostimulant is an adjuvant.

27. A vaccine according to claim 25, wherein the immunostimulant induces a predominantly Type I response.

20 28. A vaccine according to claim 25, wherein the antigen-presenting cell is a dendritic cell.

29. A method for inhibiting the development of a cancer in a patient,
25 comprising administering to a patient an effective amount of an antigen-presenting cell that expresses a polypeptide comprising at least an immunogenic portion of a lung tumor protein, or a variant thereof, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide sequence selected from the group consisting of:

30 (a) sequences recited in SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and

349;

(b) sequences that hybridize to a sequence recited in any one of SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349 under moderately stringent conditions; and

(c) complements of sequences of (i) or (ii) encoded by a polynucleotide recited in any one of SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349; and thereby inhibiting the development of a cancer in the patient.

30. A method according to claim 29, wherein the antigen-presenting cell is a dendritic cell.

31. A method according to any one of claims 21, 22 and 29, wherein the cancer is lung cancer.

32. A method for removing tumor cells from a biological sample, comprising contacting a biological sample with T cells that specifically react with a lung tumor protein, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide sequence selected from the group consisting of:

(i) polynucleotides recited in any one of SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349; and

(ii) complements of the foregoing polynucleotides; wherein the step of contacting is performed under conditions and for a time sufficient to permit the removal of cells expressing the antigen from the sample.

33. A method according to claim 32, wherein the biological sample is blood or a fraction thereof.

34. A method for inhibiting the development of a cancer in a patient, comprising administering to a patient a biological sample treated according to the method of claim 32.

35. A method for stimulating and/or expanding T cells specific for a lung tumor protein, comprising contacting T cells with at least one component selected from the group consisting of:

(a) polypeptides comprising at least an immunogenic portion of a lung tumor protein, or a variant thereof, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide sequence selected from the group consisting of:

(i) sequences recited in SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349;

(ii) sequences that hybridize to a sequence recited in any one of SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349 under moderately stringent conditions; and

(iii) complements of sequences of (i) or (ii);

(b) polynucleotides encoding a polypeptide of (a); and

(c) antigen presenting cells that express a polypeptide of (a);

under conditions and for a time sufficient to permit the stimulation and/or expansion of T cells.

36. An isolated T cell population, comprising T cells prepared according to the method of claim 35.

37. A method for inhibiting the development of a cancer in a patient, comprising administering to a patient an effective amount of a T cell population according to claim 36.

38. A method for inhibiting the development of a cancer in a patient, comprising the steps of:

(a) incubating CD4⁺ and/or CD8⁺ T cells isolated from a patient with at least one component selected from the group consisting of:

(i) polypeptides comprising at least an immunogenic portion of a lung tumor protein, or a variant thereof, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide sequence selected from the group consisting of:

10 (1) sequences recited in SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349;

(2) sequences that hybridize to a sequence recited in any one of SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349 under moderately stringent conditions; and

(3) complements of sequences of (1) or (2);

(ii) polynucleotides encoding a polypeptide of (i); and

(iii) antigen presenting cells that expresses a polypeptide of (i);

20 such that T cells proliferate; and

(b) administering to the patient an effective amount of the proliferated T cells, and thereby inhibiting the development of a cancer in the patient.

39. A method for inhibiting the development of a cancer in a patient, comprising the steps of:

(a) incubating CD4⁺ and/or CD8⁺ T cells isolated from a patient with at least one component selected from the group consisting of:

(i) polypeptides comprising at least an immunogenic portion of a lung tumor protein, or a variant thereof, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide sequence

selected from the group consisting of:

- (1) sequences recited in SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349;
- 5 (2) sequences that hybridize to a sequence recited in any one of SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349 under moderately stringent conditions; and
- (3) complements of sequences of (1) or (2);
- 10 (ii) polynucleotides encoding a polypeptide of (i); and
- (iii) antigen presenting cells that express a polypeptide of (i); such that T cells proliferate;
- (b) cloning at least one proliferated cell to provide cloned T cells; and
- 15 (c) administering to the patient an effective amount of the cloned T cells, and thereby inhibiting the development of a cancer in the patient.

40. A method for determining the presence or absence of a cancer in a patient, comprising the steps of:

- 20 (a) contacting a biological sample obtained from a patient with a binding agent that binds to a lung tumor protein, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide sequence recited in any one of SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349 or a complement of any of the
- 25 foregoing polynucleotide sequences;
- (b) detecting in the sample an amount of polypeptide that binds to the binding agent; and
- (c) comparing the amount of polypeptide to a predetermined cut-off value, and therefrom determining the presence or absence of a cancer in the patient.

41. A method according to claim 40, wherein the binding agent is an

antibody.

42. A method according to claim 43, wherein the antibody is a monoclonal antibody.

43. A method according to claim 40, wherein the cancer is lung cancer.

44. A method for monitoring the progression of a cancer in a patient, comprising the steps of:

(a) contacting a biological sample obtained from a patient at a first point in time with a binding agent that binds to a lung tumor protein, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide sequence recited in any one of SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347 and 349 or a complement of any of the foregoing polynucleotide sequences;

(b) detecting in the sample an amount of polypeptide that binds to the binding agent;

(c) repeating steps (a) and (b) using a biological sample obtained from the patient at a subsequent point in time; and

(d) comparing the amount of polypeptide detected in step (c) to the amount detected in step (b) and therefrom monitoring the progression of the cancer in the patient.

45. A method according to claim 44, wherein the binding agent is an antibody.

46. A method according to claim 45, wherein the antibody is a monoclonal antibody.

47. A method according to claim 44, wherein the cancer is a lung

cancer.

48. A method for determining the presence or absence of a cancer in a patient, comprising the steps of:

5 (a) contacting a biological sample obtained from a patient with an oligonucleotide that hybridizes to a polynucleotide that encodes a lung tumor protein, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide sequence recited in any one of SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347
10 and 349 or a complement of any of the foregoing polynucleotide sequences;

(b) detecting in the sample an amount of a polynucleotide that hybridizes to the oligonucleotide; and

(c) comparing the amount of polynucleotide that hybridizes to the oligonucleotide to a predetermined cut-off value, and therefrom determining the
15 presence or absence of a cancer in the patient.

49. A method according to claim 48, wherein the amount of polynucleotide that hybridizes to the oligonucleotide is determined using a polymerase chain reaction.

20

50. A method according to claim 48, wherein the amount of polynucleotide that hybridizes to the oligonucleotide is determined using a hybridization assay.

25

51. A method for monitoring the progression of a cancer in a patient, comprising the steps of:

(a) contacting a biological sample obtained from a patient with an oligonucleotide that hybridizes to a polynucleotide that encodes a lung tumor protein, wherein the tumor protein comprises an amino acid sequence that is encoded by a
30 polynucleotide sequence recited in any one of SEQ ID NO: 1-109, 111, 113, 115-151, 153, 154, 157, 158, 160, 162-164, 167, 168, 171, 173, 175, 177-224, 255-337, 345, 347

and 349 or a complement of any of the foregoing polynucleotide sequences;

(b) detecting in the sample an amount of a polynucleotide that hybridizes to the oligonucleotide;

(c) repeating steps (a) and (b) using a biological sample obtained from the patient at a subsequent point in time; and

(d) comparing the amount of polynucleotide detected in step (c) to the amount detected in step (b) and therefrom monitoring the progression of the cancer in the patient.

10 52. A method according to claim 51, wherein the amount of polynucleotide that hybridizes to the oligonucleotide is determined using a polymerase chain reaction.

15 53. A method according to claim 51, wherein the amount of polynucleotide that hybridizes to the oligonucleotide is determined using a hybridization assay.

54. A diagnostic kit, comprising:

(a) one or more antibodies according to claim 11; and

20 (b) a detection reagent comprising a reporter group.

55. A kit according to claim 54, wherein the antibodies are immobilized on a solid support.

25 56. A kit according to claim 54, wherein the detection reagent comprises an anti-immunoglobulin, protein G, protein A or lectin.

30 57. A kit according to claim 54, wherein the reporter group is selected from the group consisting of radioisotopes, fluorescent groups, luminescent groups, enzymes, biotin and dye particles.

58. An oligonucleotide comprising 10 to 40 contiguous nucleotides that hybridize under moderately stringent conditions to a polynucleotide that encodes a lung tumor protein, wherein the tumor protein comprises an amino acid sequence that is encoded by a polynucleotide sequence recited in any one of SEQ ID NO: 1-3, 6-8, 10-13, 15-27, 29, 30, 32, 34-49, 51, 52, 54, 55, 57-59, 61-69, 71, 73, 74, 77, 78, 80-82, 84, 86-96, 107-109, 111, 113, 125, 127, 128, 129, 131-133, 142, 144, 148-151, 153, 154, 157, 158, 160, 167, 168, 171, 173, 175, 179, 182, 184-186, 188-191, 193, 194, 198-207, 209, 210, 213, 214, 217, 220-224, 253, 254-258, 260, 262-264, 270, 272, 275, 276, 279-281, 286, 287, 291, 293, 295, 296, 300, 302, 308-310, 313, 315-317, 323, 345, 347 and 349 or a complement of any of the foregoing polynucleotides.

59. A oligonucleotide according to claim 58, wherein the oligonucleotide comprises 10-40 contiguous nucleotides recited in any one of SEQ ID NO: 1-3, 6-8, 10-13, 15-27, 29, 30, 32, 34-49, 51, 52, 54, 55, 57-59, 61-69, 71, 73, 74, 77, 78, 80-82, 84, 86-96, 107-109, 111, 113, 125, 127, 128, 129, 131-133, 142, 144, 148-151, 153, 154, 157, 158, 160, 167, 168, 171, 173, 175, 179, 182, 184-186, 188-191, 193, 194, 198-207, 209, 210, 213, 214, 217, 220-224, 253, 254-258, 260, 262-264, 270, 272, 275, 276, 279-281, 286, 287, 291, 293, 295, 296, 300, 302, 308-310, 313, 315-317, 323, 345, 347 and 349.

20

60. A diagnostic kit, comprising:

- (a) an oligonucleotide according to claim 59; and
- (b) a diagnostic reagent for use in a polymerase chain reaction or hybridization assay.

25

SEQUENCE LISTING

<110> Corixa Corporation et al.

<120> COMPOUNDS AND METHODS FOR THERAPY
AND DIAGNOSIS OF LUNG CANCER

<130> 210121.45501PC

<140> PCT

<141> 2000-04-03

<160> 350

<170> FastSEQ for Windows Version 3.0

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<223> n = A,T,C or G

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gttaatatgt ttgtaaactc atgtacagtt ttttttgagg gggaagcaat gggaanggta	240
naaattacaa atagaatcat ttgctgtaat ccttaaatgg caaacgggtca ggccacgtga	300
aaaaaaaaaa aaaaa	315

<210> 2

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<213> Homo sapien

<400> 2

atthaggtt aagattttgt ttacccttgt tactaaggag caaattagta ttaaagtata	60
atatatataa acaaatacaa aaagttttga gtggttcagc ttttttattt tttttaatgg	120
cataactttt aacaacactg ctctgtaatg ggttgaactg tggtaactcag actgagataa	180
ctgaaatgag tggatgtata gtgttattgc ataattatcc cactatgaag caaagggact	240
ggataaaattc ccagctctaga ttattagcct ttgttaacca tcaagcacct agaagaagaa	300
ttattggaaa ttttgcctc tgtaactggc actttggggg gtgacttatc ttttgccttt	360
gtaaaaaaaaa aaaaaaaaaa	380

<210> 3

<211> 346

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<220>
 <221> misc_feature
 <222> (1)...(346)
 <223> n = A,T,C or G

<400> 3
 ttgtaagtat acaatttttag aaaggattaa atgttattga tcattttact gaatactgca 60
 catcctcacc atacaccatc cactttccaa taacatttaa tcctttctaa aattgtâagt 120
 atacaattgt actttctttg gattttcata acaaataac catagactgt taattttatt 180
 gaagtttctt taatggaatg agtcattttt gtcttgtgct tttgaggta cctttgcttt 240
 gacttccaac aatttgatca tatagtgttg agctgtggaa atctttaagt ttattctata 300
 gcaataattt ctattnnnag annccngggn naaaannann annaaa 346

<210> 4
 <211> 372
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(372)
 <223> n = A,T,C or G

<400> 4
 actagtctca ttactccaga attatgctct tgtacctgtg tggctgggtt tcttagtctg 60
 tggtttggtt tggttttttg aactggtatg taggggtggtt cacagtctta atgtaagcac 120
 tcctttctcc aagtgtgctt ttgtggggac aatcattctt tgaacattag agaggaaggc 180
 agttcaagct gttgaaaaga ctattgctta tttttgtttt taaagaccta cttgacgtca 240
 tgtggacagt gcacgtgcct tacgtacat cttgttttct aggaagaagg ggatgcnggg 300
 aaggantggg tgctttgtga tggataaaac gnctaaataa cacaccttta cattttgaaa 360
 aaaacaaaac aa 372

<210> 5
 <211> 698
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(698)
 <223> n = A,T,C or G

<400> 5
 actagtanga tagaaacact gtgtcccgag agtaaggaga gaagctacta ttgattagag 60
 cctaaccag gtttaactgca agaagaggcg ggatactttc agctttccat gtaactgtat 120
 gcataaagcc aatgtagtcc agtttctaag atcatgttcc aagctaactg aatcccactt 180
 caatacacac tcatgaactc ctgatggaac aataacaggc ccaagcctgt ggtatgatgt 240
 gcacacttgc tagactcaga aaaaatacta ctctcataaa tgggtgggag tattttgggt 300
 gacaacctac tttgcttggc tgagtgaagg aatgatattc atatnttcac ttattccatg 360
 gacatttagt tagtgctttt tatataccag gcatgatgct gagtgacact cttgtgtata 420
 tntccaaatn ttngtncngt cgctgcacat atctgaaatc ctatattaag antttcccaa 480
 natgangtcc ctggtttttc cacgccactt gatcngtcaa ngatctcacc tctgtntgtc 540
 ctaaaacnt ctncnnang gttagacngg acctctcttc tcccttcccg aanaatnaag 600
 tgtgngaaga nancncnch cccccctnch tncnnctng ccngctnnnc cnctgtngg 660

gggngcggcc cccgcggggg gacccccccn ttttcccc

698

<210> 6
 <211> 740
 <212> DNA
 <213> Homo sapien

 <220>
 <221> misc_feature
 <222> (1)...(740)
 <223> n = A,T,C or G

<400> 6
 actagtcaaa aatgctaaaa taatttgga gaaaatattt ttttaagtagt gttatagttt 60
 catgtttatc ttttattatg tnttgtgaag ttgtgtcttt tcaactaatta cctatactat 120
 gccaatattt ccttatatct atccataaca tttatactac atttgtaaga gaatatgcac 180
 gtgaaactta acactttata aggtaaaaat gaggtttcca agatttaata atctgatcaa 240
 gttcttggtta tttccaaata gaatggactt ggtctgttaa ggggctaagg gagaagaaga 300
 agataagggtt aaaagtgtgtt aatgacccaaa cattctaaaa gaaatgcaaa aaaaaattta 360
 ttttcaagcc ttggaactat ttaaggaaag caaatcatt tcctanatgc atatcatttg 420
 tgagantttc tcantaatat cctgaatcat tcatttcagc tnaggcttca tgttgactcg 480
 atatgtcatc tagggaaagt ctatttcattg gtccaaacct gttgccatag ttggttaggc 540
 tttcctttaa ntgtgaanta ttnacangaa atttctctt tnanagttct tnatagggtt 600
 aggggtgtgg gaaaagcttc taacaatctg tagtgttncg tggtatctgt ncagaaccan 660
 aatnacggat cgnangaagg actgggtcta tttacangaa cgaatnatct ngttmmtgt 720
 gtnnncaact ccngggagcc 740

<210> 7
 <211> 670
 <212> DNA
 <213> Homo sapien

 <220>
 <221> misc_feature
 <222> (1)...(670)
 <223> n = A,T,C or G

<400> 7
 gctggggagc tcggcatggc ggtccccgct gcagccatgg ggcctcggc gttgggccag 60
 agcgcccccg gctcgatggc cccgtggtgc tcagtgaaga gcggccccgc gcgctacgtg 120
 cttgggatgc aggagctgtt ccggggccac agcaagaccg cgagttcctg gcgcacagcg 180
 ccaaggtgca ctcggtggcc tggagttgag acgggcgtcg cctacctcgg ggtcttcgac 240
 aagacgccac gtcttcttgc tgganaanga ccgttggtca aagaaaacaa ttatcgggga 300
 catggggata gtgtggacca ctttgttggc atccaagtaa tcctgacctt tttgttacgg 360
 cgtctggaga taaaaccatt cgcattctggg atgtgaggac tacaaaatgc attgccactg 420
 tgaactactaa aggggagaac attaatatct gctggantcc tgatgggcan accattgctg 480
 tagcnacaag gatgatgtgg tgactttatt gatgccaaaga aacccccgtc caaagcaaaa 540
 aaacanttcc aanttcgaag tcaccnaaat ctcttggaac aatgaacatn aatatnttct 600
 tcctgacaat ggnccctggg tgnntcacat cctcagctnc cccaaaactg aancctgtnc 660
 natccacccc 670

<210> 8
 <211> 689
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (689)
 <223> n = A,T,C or G

<400> 8
 actagtatct aggaatgaac agtaaaagag gagcagttgg ctacttgatt acaacagagt 60
 aaatgaagta ctggatttgg gaaaacctgg ttttattaga acatatggaa tgaaagccta 120
 cacctagcat tgcctactta gcccctgaa ttaacagagc ccaattgaga caaacccctg 180
 gcaacaggaa attcaaggga gaaaaagtaa gcaacttggg ctaggatgag ctgactccct 240
 tagagcaaag ganagacagc ccccattacc aaataccatt tttgcctggg gcttgtgcag 300
 ctggcagtgt tcctgcccga gcatggcacc ttatngtttt gatagcaact tcgttgaatt 360
 ttcaccaact tattacttga aattataata tagcctgtcc gtttgcctgn tccaggctgt 420
 gatatatntt cctagtgggt tgactttnaa aataaatnag gtttantttt cccccccnn 480
 cnnctnctnc nntcnctcnn cnncccccc cncctngtcc tccnnnnntn gggggggcnn 540
 cccccnoggg ggacccccct ttggtccctt agtggagggt natggcccc ggnnttatcc 600
 nggcctann tttccccgtn mnaaatgntt cccctccca ntccnccac ctcaanccgg 660
 aagcctaagt ttntaccctg ggggtcccc 689

<210> 9
 <211> 674
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (674)
 <223> n = A,T,C or G

<400> 9
 gtccactctc ctttgagtgt actgtcttac tgtgcactct gtttttcaac tttctagata 60
 taataaatgc ttgttctata gtggagtaag agctcacaca cccaaggcag caagataact 120
 gaaaaaagcg aggctttttt gccaccttgg taaaggccag ttcactgcta tagaactgct 180
 ataagcctga aggggaagtag ctatgagact ttccattttt cttagtctc ccaataggct 240
 ccttcatgga aaaaggcttc ctgtaataat tttcacctaa tgaattagca gtgtgattat 300
 ttctgaaata agagacaaat tgggcgcag agtcttctctg tgatttaaaa taacaaccc 360
 aaagttttgt ttggtcttca ccaaaggaca tactctaggg ggtatgttgt tgaagacatt 420
 caaaaacatt agctgttctg tctttcaatt tcaagttatt ttggagactg cctccatgtg 480
 agttaattac tttgctctgg aactagcatt attgtcatta tcatcacatt ctgtcatcat 540
 catctgaata atattgtgga tttccccctc tgcttgcatc ttcttttgac tctctggga 600
 anaaatgtca aaaaaaagg tcgatctact cngcaaggnc catctaata caatgcgtgga 660
 aggaccnct gcc 674

<210> 10
 <211> 346
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (346)
 <223> n = A,T,C or G

<400> 10

actagtctgc	tgatagaaag	cactatacat	cctattgttt	ctttctttcc	aaaatcagcc	60
ttctgtctgt	aacaaaaatg	tactttatag	agatggagga	aaaggtctaa	tactacatag	120
ccttaagtgt	ttctgtcatt	gttcaagtgt	atcttctgta	acagaaacat	atttggaatg	180
tttttctttt	ccccttataa	attgtaattc	ctgaaatact	gctgctttta	aaagtcctac	240
tgtcagatta	tattatctaa	caattgaata	ttgtaaatat	acttgtctta	cctctcaata	300
aaaggtgact	tttctattan	nnagnngnnn	gnnnnataaa	anaaaa		346

<210> 11

<211> 602

<212> DNA

<213> Homo sapien

<400> 11

actagtaaaa	agcagcattg	ccaaataatc	cctaattttc	cactaaaaat	ataatgaaat	60
gatgttaagc	tttttgaaaa	gtttagggtta	aacctactgt	tgtagatta	atgtatttgt	120
tgcttccctt	tatctggaat	gtggcattag	cttttttatt	ttaaccctct	ttaattotta	180
ttcaattcca	tgacttaagg	ttggagagct	aaacactggg	atctttggat	aacagactga	240
cagttttgca	taattataat	cggcattgta	catagaaagg	atatggctac	cttttggtta	300
atctgcactt	tctaaatata	aaaaaaggga	aatgaagtta	taaatcaatt	ttgtataat	360
ctgtttgaaa	catgagtgtt	atttgcttaa	tattagggct	ttgccccttt	tctgtaagtc	420
tcttgggata	ctgtgtagaa	ctgttctcat	taaacaccaa	acagttaagt	ccattctctg	480
gtactagcta	caaattcggg	ttcatattct	acttaacaat	ttaaataaac	tgaaatattt	540
ctagatgggtc	tacttctgtt	catataaaaa	caaaacttga	tttccaaaaa	aaaaaaaaaa	600
aa						602

<210> 12

<211> 685

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(685)

<223> n = A,T,C or G

<400> 12

actagtctctg	tgaaagtaca	actgaaggca	gaaagtgtta	ggattttgca	tctaattgtc	60
attatcatgg	tattgatgga	cctaagaaaa	taaaaattag	actaagcccc	caaataagct	120
gcatgcattt	gtaacatgat	tagtagattt	gaatatatag	atgtagtatn	ttgggtatct	180
aggtgtttta	tcattatgta	aaggaattaa	agtaaaggac	ttttagtttg	tttttattaa	240
atatgcatat	agtagagtgc	aaaaatatag	caaaaatana	aactaaagggt	agaaaagcat	300
tttagatatg	ccttaatnta	nnaactgtgc	caggtggccc	tcggaataga	tgccaggcag	360
agaccagtgc	ctgggtggtg	cctccccttg	tctgcccccc	tgaagaactt	ccctcacgtg	420
angtagtgcc	ctcgtagggtg	tcacgtggan	tantggganc	aggccgnncn	gtnanaagaa	480
ancanngtga	nagtttcncc	gtngangcng	aactgtccct	gngccnnnac	gctcccanaa	540
cntntccaat	ngacaatcga	gtttccnnnc	tcengnaacc	tngccgnnnn	cnngcccnnc	600
cantntgnta	accccgcgcc	cggatcgctc	tcnnntcggt	ctcncncnaa	ngggntttcn	660
cnnccgcggt	cncnnccccg	cnncc				685

<210> 13

<211> 694

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature
 <222> (1) ... (694)
 <223> n = A,T,C or G

<400> 13

cactagtcac	tcattagcgt	tttcaatagg	gctcttaagt	ccagtagatt	acgggtagtc	60
agttgacgaa	gatctgggtt	acaagaacta	attaaatgtt	tcattgcatt	tttgtaagaa	120
cagaataaatt	ttataaaatg	ttttagagtt	ataattgccg	aaaataattt	aaagacactt	180
tttctctgtg	tgtgcaaagt	tgtgtttgtg	atccattttt	tttttttttt	taggacacct	240
gtttactagc	tagctttaca	atatgccaaa	aaaggatttc	tccttgacct	catcogtggg	300
tcacctcttt	ttcccccat	gctttttgcc	ctagtttata	acaaagggaat	gatgatgatt	360
taaaaagtag	ttctgtatct	tcagtatctt	ggtcttcag	aacctctctg	ttgggaagg	420
gatcattttt	tactggtcat	ttccctttgg	agtgtactac	tttaacagat	ggaaagaact	480
cattggccat	ggaaacagcc	gangtggttg	gagccagcag	tgcatggcac	cgtccggcat	540
ctggcntgat	tggctctggc	gccgtcattg	tcagcacagt	gccatgggac	atggggaana	600
ctgactgcac	ngccaatggg	tttcatgaag	aatacngcat	ncncngtgat	cacgtnancc	660
angacgctat	gggggncana	gggcanttg	cttc			694

<210> 14
 <211> 679
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (679)
 <223> n = A,T,C or G

<400> 14

cagcgcctg	catctgtatc	cagcgcang	tcctgcagc	cccagctg	cgcgcccc	60
agtcccgac	cgttcggcc	cangctnagt	tagnctcac	catnccggtc	aaaggangca	120
ccaagtgc	caaatacctg	cngtnoggat	ntaaattcat	cttctggctt	gccgggattg	180
ctgtccntgc	cattggacta	nggtccgat	ncgactctca	gaccanganc	atcttcganc	240
naganactaa	tnatnatnt	tcagcttct	acacaggagt	ctatattctg	atcggtccg	300
gcncctcnt	gatgctggtg	ggcttcctga	gctgctg	ggctgtgcaa	gagtcctant	360
gcatgctggg	actgttcttc	ggcttcntct	tggtgatatn	cgccattgaa	atacctgcgg	420
ccatctgggg	atattccact	ncgatnatgt	gattaaggaa	ntccacggag	ttttacaagg	480
acacgtacaa	cnacctgaaa	accnnggatg	anccccaccg	ggaancnctg	aangccatcc	540
actatgcgtt	gaactgcaat	ggtttggctg	gggnccttga	acaattta	cncatacatc	600
tgccccann	aaaggacntn	ctcgannct	tcnccgtgna	attcngttct	gatnccatca	660
cagaagtctc	gaacaatcc					679

<210> 15
 <211> 695
 <212> DNA
 <213> Homo sapien

<220>
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 <222> (1) ... (695)
 <223> n = A,T,C or G

<400> 15

actagtggat	aaaggccagg	gatgctgctc	aacctcctac	catgtacagg	gacgtctccc	60
cattacaact	acccaatccg	aagtgtcaac	tgtgtcagga	ctaanaaacc	ctggttttga	120

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ttaaaaaagg gcctgaaaaa aggggagcca caaatctgtc tgcttccctca cnttantcnt 180
tggcaaatna gcattctgtc tcnttggtcg cngcctcanc ncaaaaaanc ngaactcnat 240
cnggcccagg aatacatctc ncaatnaacn aaattganca aggcnnntggg aaatgccnga 300
tgggattatc ntccgcttgt tgancttcta agtttcnttc ccttcattcn accctgccag 360
ccnagttctg ttagaaaaat gccngaattc naacnccggt tttcntactc ngaatttaga 420
tctncanaaa ctctctggcc acnattcnaa ttanngnca cgnacanatn ccttccatna 480
ancncacccc acntttgana gccangacaa tgactgcntn aantgaaggc ntgaaggaaan 540
aaactttgaaa ggaaaaaaa ctttggttcc ggcccttcc aacncttctg tgttnancac 600
tgccctctng naaccctgga agcccnnga cagtgttaca tgttggtcta nnaaacngac 660
ncttnaatnt cnatcttccc nanaacgatt ncnc 695

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<210> 16

<211> 669

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (669)

<223> n = A,T,C or G

<400> 16

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cgccgaagca gcagcgagg ttgtccccgt tccccctccc ccttcccttc tccgggtgcc 60
tccccgggcc ccttacctc cacagtcccg gtcccgccat gtcccagaaa caagaagaag 120
agaaccctgc ggaggagacc ggcgaggaga agcaggacac gcaggagaaa gaaggtattc 180
tgccctgagag agctgaagag gcaaagctaa aggccaaata cccaagccta ggacaaaagc 240
ctggaggctc cgacttcctc atgaagagac tccagaaagg gcaaaagtac tttgactcng 300
gagactacaa catggccaaa gccaacatga agaataagca gctgccaaagt gcangaccag 360
acaagaacct ggtgactggt gatcacatcc ccacccaca ggatctgccc agagaaagtc 420
ctcgctcgtc accagcaagc ttgcggttg ccaagttgaa tgatgctgcc ggggctctgc 480
canatctgag acgcttcctt ccttgcctca cccgggtcct gtgctggctc ctgcccttcc 540
tgcttttgca gccangggc aggaagtggc ncnggtngtg gctggaaagc aaaacccttt 600
cctgttggtg tcccacccat ggagcccctg gggcgagccc angaacttga nccttttgt 660
tntcttncc 695

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<210> 17

<211> 697

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (697)

<223> n = A,T,C or G

<400> 17

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gcaagatatg gacaactaag tgagaaggta atnctctact gctctagntn ctccnggcnn 60
gacgcgtga ggagannnac gctggcccan ctgcggcca cacacgggga tcntggtnat 120
gcctgccan gggancccca ncnetcgan cccatntcac acccgnnccn tncgcccacn 180
ncctggctcn cncngcccng nccagctcnc gnceccctec gccnnnetcn ttrnctctc 240
cncnccctec ncnacnact cctaccnng gctccctccc cagccccccc ccgcaanct 300
ccacnacncc ntennncga ancnecnetc gcnctcngcc cngccccct gcccccgcc 360
cncnacnncg cgncccccg cgcncgcngc ctncccccct cccacnacag ncncacccgc 420
agncaegcnc tccgcccct gacgcccnn cccgcgcgc tcaccttcat ggncnacng 480
ccccgctcnc ncnctgcnc gccgncngg cgcgccccc cncnccngtn ccncnngng 540

```

```

ccccngcngn angcngtgcg cnncaangncc gngccggnncn ncaccctccg nccnccgccc 600
cgcccgctgg gggctcccgc cncgcggntc antcccccnc cntnccgcca ctntccgntc 660
cnnctctcnc gctcngcgcn cgcccnccnc cccccc 697

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<210> 18
<211> 670
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)... (670)
<223> n = A,T,C or G

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<400> 18
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gggacggctg cccgcggggc cccggggcat gggcacggcc ctgaagctgt tgctgggggc 180
cggcgccgtg gcctacgggt tgcgcgaatc tgtgttcacc gtggaaggcg ggcncagagc 240
catcttcttc aatcgggatc gtggagtgc caggacacta tcttggggccg anggccttca 300
cttcaggatc cttggttcca gtaccccanc atctatgaca ttcggggccag acctcgaaaa 360
aatctcctcc ctacaggctc caaagacctc cagatggtga atatctccct gcgagtgttg 420
tctcgaccaa tgctcangaa cttcctaaca tgttccancg cctaagggct ggactacnaa 480
gaacgantgt tgccgtccat tgtcacgaag tgctcaagaa tttnggtggc caagttcaat 540
gncctcacnn ctgatcnccc agcggggcca agttanccct ggttgatccc cgggganctg 600
acnnaaaagg gccaaaggact tcccctcatc ctggataatg tggcctcac aaagctcaac 660
tttanccacc 670

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<210> 19
<211> 606
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)... (606)
<223> n = A,T,C or G

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<400> 19
actagtgcc aacctagctc ccaggccagt tctctgaatg tcgaggagtt ccaggatctc 60
tggcctcagt tgtccttggt tattgatggg ggacaaattg gggatggcca gagccccgag 120
tgtcgcttg gctcaactgt ggttgatttg tctgtgcccg gaaagtttg catcattcgt 180
ccaggtctg cctggaaag tactacagcc atcctccaac agaagtacgg actgctcccc 240
tcacatgcgt cctacctgtg aaactctggg aagcaggaag gcccaagacc tgggtgctgga 300
tactatgtgt ctgtccactg acgactgtca aggcctcatt tgcagaggcc accggagcta 360
gggcactagc ctgactttta aggcagtgtg tctttctgag cactgtagac caagcccttg 420
gagctgctgg tttagccttg cacctgggga aaggatgtat ttatttgtat tttcatatat 480
cagccaaaag ctgaatggaa aagttnagaa cattcctagg tggccttatt ctaataagtt 540
tcttctgtct gttttgtttt tcaattgaaa agttattaaa taacagattt agaatctagt 600
gagacc 606

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<210> 20
<211> 449
<212> DNA
<213> Homo sapien

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<400> 20
 actagtaaac aacagcagca gaaacatcag tatcagcagc gtcgccagca ggagaatatg 60
 cagcgccaga gccgaggaga acccccgctc cctgaggagg acctgtccaa actcttcaaa 120
 ccaccacagc cgcctgccag gatggactcg ctgctcattg caggccagat aaacacttac 180
 tgccagaaca tcaaggagtt cactgccc aaacttaggca agctcttcatt ggcccaggct 240
 cttcaagaat acaacaacta agaaaaggaa gtttccagaa aagaagttaa catgaactct 300
 tgaagtcaca ccagggaac tcttggaaga aatatatttg catattgaaa agcacagagg 360
 atttcttttag tgtcattgcc gattttggct ataacagtgt ctttctagcc ataataaaat 420
 aaaacaaaat cttgactgct tgctcaaaa 449

<210> 21
 <211> 409
 <212> DNA
 <213> Homo sapien

<400> 21
 tatcaatcaa ctggtgaata attaaacaat gtgtggtgtg atcatacaaa ggttaccact 60
 caatgataaa aggaacaagc tgcctatatg tggaacaaca tggatgcatt tcagaaactt 120
 tatgttgagt gaaagaacaa acacggagaa catactatgt ggttctcttt atgtaacatt 180
 acagaaataa aaacagaggc aaccaccttt gaggcagtat ggagtgagat agactggaaa 240
 aaggaaggaa ggaaactcta cgctgatgga aatgtctgtg tcttcattgg gtggtagtta 300
 tgtggggata tacatttgtc aaaattttatt gaactatata ctaaagaact ctgcatttta 360
 ttgggatgta aataatacct caattaaaaa gacaaaaaaa aaaaaaaaaa 409

<210> 22
 <211> 649
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (649)
 <223> n = A,T,C or G

<400> 22
 acaattttca ttatcttaag cacattgtac atttctacag aacctgtgat tattctcgca 60
 tgataaggat ggtacttgca tatggtgaat tactactgtt gacagtttcc gcagaaatcc 120
 tatttcagtg gaccaacatt gtggcatggc agcaaatgcc aacattttgt ggaatagcag 180
 caaatctaca agagaccctg gttggttttt cgttttgttt tctttgtttt tcccccttc 240
 tcctgaatca gcagggatgg aangagggta gggaagtat gaattactcc ttccagtagt 300
 agctctgaag tgtcacattt aatatcagtt ttttttaaac atgattctag ttnaatgtag 360
 aagagagaag aaagaggaag tgttcaactt tttaatacac tgatttagaa atttgatgtc 420
 ttatatcagt agttctgagg tattgatagc ttgctttatt tctgccttta cgttgacagt 480
 gttgaagcag ggtgaataac taggggcata tatatttttt ttttttgtaa gctgtttcat 540
 gatgttttct ttggaatttc cggataagtt caggaaaaca tctgcatgtt gttatctagt 600
 ctgaagttcn tatccatctc attacaacaa aaacnccag aacggnntg 649

<210> 23
 <211> 669
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature

<222> (1)...(669)

<223> n = A,T,C or G

<400> 23

actagtgcg	tactggctga	aatccctgca	ggaccaggaa	gagaaccagt	tcagactttg	60
tactctcagt	caccagctct	ggaattagat	aaattccttg	aagatgtcag	gaatgggatc	120
tatcctctga	cagcctttgg	gctgcctcgg	ccccagcagc	cacagcagga	ggaggtgaca	180
tcacctgtcg	tgccccctc	tgtcaagact	ccgacacctg	aaccagctga	ggtggagact	240
cgcaaggtgg	tgctgatgca	gtgcaacatt	gagtccggtg	aggaggaggt	caaacaccac	300
ctgacacttc	tgctgaagtt	ggaggacaaa	ctgaaccggc	acctgagctg	tgacctgatg	360
ccaaatgaga	atatccccga	gttggcggct	gagctgggtg	agctgggctt	cattagttag	420
gctgaccaga	gccggttgac	ttctctgcta	gaagagactt	gaacaagttc	aattttgcca	480
ggaacagtac	cctcaactca	gccgctgtca	ccgtctcctc	ttagagctca	ctcgggccag	540
gccctgatct	gcgctgtggc	tgtcctggac	gtgctgcacc	ctctgtcctt	ccccccagtc	600
agtattacct	gtgaagccct	tccctccttt	attattcagg	anggctgggg	gggctccttg	660
nttctaacc						669

<210> 24

<211> 442

<212> DNA

<213> Homo sapien

<400> 24

actagtacca	tcttgacaga	ggatacatgc	tcccaaaacg	tttgttacca	cacttaaaaa	60
tactgcat	cattaagcat	cagtttcaaa	attatagcca	ttcatgattt	actttttcca	120
gatgactatc	attattctag	tcctttgaat	ttgtaagggg	aaaaaaaaa	aaaacaaaaa	180
cttacgatgc	acttttctcc	agcacatcag	atttcaaatt	gaaaattaaa	gacatgctat	240
ggtaatgcac	ttgctagtac	tacacacttt	ggtacaacaa	aaaacagagg	caagaaacaa	300
cggaaagaga	aaagccttcc	tttgttggcc	cttaaaactga	gtcaagatct	gaaatgtaga	360
gatgatctct	gacgatacct	gtatgttctt	attgtgtaaa	taaaattgct	ggtatgaaat	420
gacctaataa	aaaaaaaaa	aa				442

<210> 25

<211> 656

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(656)

<223> n = A,T,C or G

<400> 25

tgcaagtacc	acacactggt	tgaattttgc	acaaaaagtg	actgtaggat	caggtgatag	60
ccccggaatg	tacagtgtct	tggtgcacca	agatgccttc	taaaggctga	cataccttgg	120
accctaattg	ggcagagagt	atagccctag	cccagtggtg	acatgaccac	tccttttggg	180
aggcctgagg	tagaggggag	tggtatgtgt	tttctcagtg	gaagcagcac	atgagtgggt	240
gacaggatgt	tagataaagg	ctctagttag	ggtgtcattg	tcatttgaga	gactgacaca	300
ctcctagcag	ctggtaaaag	ggtgctggan	gccatggagg	anctctagaa	acattagcat	360
gggctgatct	gattacttcc	tggtatcccg	ctcactttta	tggaagtct	tattagangg	420
atgggacagt	tttccatata	cttgctgtgg	agctctggaa	cactctctaa	atttcctct	480
attaaaaatc	actgccctaa	ctacacttcc	tccttgaagg	aatagaaatg	gaactttctc	540
tgacatantt	cttggcatgg	ggagccagcc	acaaatgana	atctgaacgt	gtccaggttt	600
ctcctganac	tcattctacat	agaattgggt	aaaccctccc	ttggaataag	gaaaaa	656

<210> 26
 <211> 434
 <212> DNA
 <213> Homo sapien
 <220>
 <221> misc_feature
 <222> (1)...(434)
 <223> n = A,T,C or G

<400> 26
 actagtctcag actgccacgc caaccccaga aaatacccca catgccagaa aagtgaagtc 60
 ctagggtgtt ccatctatgt ttcaatctgt ccatctacca ggcctcgcga taaaaacaaa 120
 acaaaaaaac gctgccaggt tttagaagca gttctggtct caaaaccatc aggatcctgc 180
 caccagggtt cttttgaaat agtaccacat gtaaaagggg atttggcttt cacttcatct 240
 aataactgaa ttgtcaggct ttgattgata attgtagaaa taagtagcct tctgttgtgg 300
 gaataagtta taatcagtat tcatctcttt gttttttgtc actcttttct ctctaattgt 360
 gtcatttgta ctggttgaaa aatatttctt ctatnaaatt aaactaacct gccttaaaaa 420
 aaaaaaaaaa aaaa 434

<210> 27
 <211> 654
 <212> DNA
 <213> Homo sapien
 <220>
 <221> misc_feature
 <222> (1)...(654)
 <223> n = A,T,C or G

<400> 27
 actagtccaa cacagtcaga aacattgttt tgaatcctct gtaaaccaag gcattaatct 60
 taataaacca ggatccattt aggtaccact tgatataaaa aggatatcca taatgaatat 120
 tttatactgc atcctttaca ttagccacta aatacgttat tgcttgatga agacctttca 180
 cagaatccta tggattgcag catttcactt ggctacttca taccatgcc ttaaagaggg 240
 gcagtttctc aaaagcagaa acatgccgcc agttctcaag ttttctcct aactccattt 300
 gaatgtaagg gcagctggcc cccaatgtgg ggaggtccga acattttctg aattccatt 360
 ttcttgttcg cggctaaatg acagtttctg tcattactta gattccgatc tttcccaag 420
 gtgttgattt acaaagaggc cagctaatag cagaaatcat gacctgaaa gagagatgaa 480
 attcaagctg tgagccaggc agganctcag tatggcaaag gtcttgagaa tcngccattt 540
 ggtacaaaaa aaatttttaa gcntttatgt tataccatgg aaccatagaa anggcaaggg 600
 aattgttaag aanaatttta agtgtccaga ccanaanga aaaaaaaaaa aaaa 654

<210> 28
 <211> 670
 <212> DNA
 <213> Homo sapien
 <220>
 <221> misc_feature
 <222> (1)...(670)
 <223> n = A,T,C or G

<400> 28
 cgtgtgcaca tactgggagg atttccacag ctgcacggtc acagccctta cggattgcc 60

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ggaaggggag aaagatatgt gggataaact gagaaaagaa nccaaaaacc tcaacatcca 120
aggcagctta ttcgaactct ggggcagcgg caacggggcg gcgggggtccc tgctcccggc 180
gttcccggtg ctctcgtgtg ctctctcggc agcttttagcg acctgncctt ccttctgagc 240
gtggggccag cccccccgc ggcgcacc cactctcact ccatgctccc ggaaatcgag 300
aggaagatca ttagttcttt ggggacgttn gtgattctct gtgatgctga aaaacactca 360
tatagggaat gtgggaaatc ctganctctt tnttatntcg tntgatttct tgtgttttat 420
ttgccaaaat gttaccaatc agtgaccaac cnagcacagc caaaaatcgg acntcngctt 480
tagtccgtct tcacacacag aataagaaaa cggcaaacc accccacttt tnantttnat 540
tattactaan ttttttctgt tgggcaaaag aatctcagga acngccctgg ggccnccgta 600
ctanagttaa ccnagctagt tncatgaaaa atgatgggct ccncctcaat gggaaagcca 660
agaaaaagnc

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<210> 29
<211> 551
<212> DNA
<213> Homo sapien
<220>
<221> misc_feature
<222> (1) ... (551)
<223> n = A,T,C or G

```

```

<400> 29
actagtcttc cacagcctgt gaatccccct agacctttca agcatagtga gcggagaaga 60
agatctcagc gtttagccac cttaccatg cctgatgatt ctgtagaaaa ggtttcttct 120
ccctctccag ccactgatgg gaaagtattc tccatcagtt ctcaaatca gcaagaatct 180
tcagtaccag aggtgcctga tgttgacat ttgccacttg agaagctggg accctgtctc 240
cctcttgact taagtcgtgg ttcagaagtt acagcacgg tagcctcaga ttctctttac 300
cgtaatgaat gtcccagggc agaaaaagag gatacncaga tgcttccaaa tccttcttcc 360
aaagcaatag ctgatgggaa gaggagctcc agcagcagca ggaatatcga aaacagaaaa 420
aaaagtgaat ttgggaagac aaaagctcaa cagcatttgg taaggagaaa aganaagatg 480
aggaaggaag agagaagaga gacnaagatc nctacggacc gnnncggaag aagaagaagn 540
aaaaaanaaa a 551

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<210> 30
<211> 684
<212> DNA
<213> Homo sapien
<220>
<221> misc_feature
<222> (1) ... (684)
<223> n = A,T,C or G

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<400> 30
actagttcta tctggaaaaa gccggggttg gaagaagctg tggagagtgc gtgtgcaatg 60
cgagactcat ttcttggaag catccctggc aaaaatgcag ctgagtacaa ggttatcact 120
gtgatagaac ctggactgct ttttgagata atagagatgc tgagctctga agagacttcc 180
agcacctctc agttgaatga attaataatg gcttctgagt caactttact ggctcaggaa 240
ccacgagaga tgactgcaga tgtaatcgag cttaaaggga aattcctcat caacttagaa 300
ggtggtgata ttcgtgaaga gtcttcctat aaagtaattg tcatgccgac tacgaaagaa 360
aaatgcccc gttgttgga gtatacagcg ggagtcttca gatacactgt gtcctcgatg 420
tgcagaagtt gtcagtggga aaatagtatt aacagctcac tcgagcaaga accctcctga 480
cagtactggg ctagaagttt ggatggatta tttacaatat aggaagaaa gccaagaatt 540
aggtnatgag tggatgagta aatggtggan gatggggaat tcaaatcaga attatggaag 600

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aagttnttcc tgttactata gaaaggaatt atgtttatattt acatgcagaa aatatanatg 660
 tgtggtgtgt accgtggatg gaan 684

<210> 31
 <211> 654
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (654)
 <223> n = A,T,C or G

<400> 31

ggcgcagaaaa ggaaccaata tttcagaaac aagcttaata ggaacagctg cctgtacatc	60
aacatcttct cagaatgacc cagaagttat catcgtggga gctggcgtgc ttggctctgc	120
tttggcagct gtgctttcca gagatggaag aaaggtgaca gtcattgaga gagacttaaa	180
agagcctgac agaatagttg gagaattcct gcagccgggt gggtatcatg ttctcaaaga	240
ccttggctctt ggagatacag tgggaaggtct tgatgccag gttgtaaatg gttacatgat	300
tcacatgacag ggaaagcaaa tcagangttc agattcctta ccctctgtca gaaaacaatc	360
aagtgcagag tggaaagagct ttccatcacg gaagattcat catgagtctc cggaaagcag	420
ctatggcaga gcccaatgca aagtttattg aaggtgttgt gttacagtta ttagaggaag	480
atgatgttgt gatggagtt cagtacaagg ataaagagac tgggagatat caaggaactc	540
catgctccac tgactgttgt tgcagatggg cttttctcca anttcaggaa aagcctggtc	600
tcaataaagt ttctgtatca ctcatgtgtg tggcttctta tgaagaatgc nccc	654

<210> 32
 <211> 673
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (673)
 <223> n = A,T,C or G

<400> 32

actagtgaag aaaaagaaat tctgatacgg gacaaaaatg ctcttcaaaa catcattctt	60
tatcacctga caccaggagt tttcattgga aaaggatttg aacctggtgt tactaacatt	120
ttaaagacca cacaaggaag caaaatcttt ctgaaagaag taaatgatac acttctggtg	180
aatgaattga aatcaaaaga atctgacatc atgacaacaa atggtgtaat tcatgttgta	240
gataaactcc tctatccagc agacacacct gttggaaatg atcaactgct ggaaatactt	300
aataaattaa tcaaatacat ccaaattaag tttgttcgtg gtagcacctt caaagaaatc	360
cccgtagctg tctatnagcc aattattaaa aaatacacca aaatcattga tgggagtgcc	420
tgtgggaaat aactgaaaaa gagaccgaga agaacgaatc attacagggtc ctgaaataaa	480
atacctagga tttctactgg aggtggagaa acagaagaac tctgaagaaa ttgttacaag	540
aagangtccc aaggtcacca aattcattga aggtggtgat ggtctttatt tgaagatgaa	600
gaaattaaaa gacgcttcag ggagacnccc catgaaggaa ttgccagcca caaaaaatt	660
cagggattag aaa	673

<210> 33
 <211> 673
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (673)
 <223> n = A,T,C or G

<400> 33

actagtttatt	tacttttctc	cgcttcagaa	ggtttttcag	actgagagcc	taagcatact	60
ggatctgttg	tttcttttgg	gtctcacctc	atcagtgtgc	atagtggcag	aaattataaa	120
gaagggttgaa	aggagcaggg	aaaagatcca	gaagcatggt	agttcgacat	catcatcttt	180
tcttgaagta	tgatgcatat	tgcattatth	tatttgcaaa	ctaggaattg	cagtctgagg	240
atcatttaga	agggcaagtt	caagaggata	tgaagatttg	agaacttttt	aactattcat	300
tgactaaaaa	tgaacattaa	tgttnaagac	ttaagacttt	aacctgctgg	cagtcccaaa	360
tgaattatg	caactttgat	atcatattcc	ttgatttaaa	ttgggctttt	gtgattgant	420
gaaactttat	aaagcatatg	gtcagttatt	tnattaaaaa	ggcaaaacct	gaaccacctt	480
ctgcacttaa	agaagtctaa	cagtacaaat	acctatctat	cttagatgga	tntatttntt	540
tntattttta	aatattgtac	tatttatggt	nggtggggct	ttcttactaa	tacacaaatn	600
aatttatcat	ttcaanggca	ttctatttgg	gtttagaagt	tgattccaag	nantgcatat	660
ttcgctactg	tnt					673

<210> 34
 <211> 684
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (684)
 <223> n = A,T,C or G

<400> 34

actagttttat	tcaagaaaag	aactttactga	ttcctctgtt	cctaaagcaa	gagtggcagg	60
tgatcagggc	tggtgtagca	tccggttcct	ttagtgcagc	taactgcatt	tgctactgat	120
gaccaaggag	gaaatcacta	agacatttga	gaagcagtg	tatgaacgtt	cttggaacaag	180
ccacagttct	gagccttaac	cctgtagttt	gcacacaaga	acgagctcca	cctccccctc	240
ttcaggagga	atctgtgcgg	atagattggc	tggacttttc	aatggttctg	ggttgcaagt	300
gggcactggt	atggctgggt	atggagcgga	cagccccagg	aatcagagcc	tcagcccggc	360
tgcttggtt	gaaggtacag	gtgttcagca	ccttcggaaa	aagggcataa	agtngtgggg	420
gacaattctc	agtccaagaa	gaatgcattg	accattgctg	gctatttgct	tncttagtan	480
gaattggatn	catttttgac	cangatnntt	ctnctatgct	tnnttgcaat	gaaatcaa	540
ccgcattat	ctacaagtgg	tatgaagtcc	tgcncccccc	agagaggctg	ttcaggcnat	600
gtcttccaag	ggcaggggtg	gttacaccat	ttacctccc	ctctcccccc	agattatgna	660
cncagaagga	atttntttcc	tccc				684

<210> 35
 <211> 614
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (614)
 <223> n = A,T,C or G

<400> 35

actagtccaa	cgcgttngcn	aatattcccc	tggtagccta	cttccttacc	ccogaatatt	60
------------	------------	------------	------------	------------	------------	----

ggtaagatcg	agcaatggct	tcaggacatg	ggttctcttc	tcctgtgac	attcaagtgc	120
tcaactgcag	aagactggct	tgtctcagtg	tntcaacctc	accagggtg	tctcttggtc	180
cacacctcgc	tccctgttag	tgccgtatga	cagcccccat	canatgacct	tggccaagtc	240
acggtttctc	tgtggtcaat	gttggtnggc	tgattggtgg	aaagtanggt	ggaccaaagg	300
aagncncgtg	agcagncanc	nccagttctg	caccagcagc	gcctccgtcc	tactngggtg	360
ttccngtttc	tcctggccct	gngtgggcta	nggcctgatt	cggggaanatg	cctttgcang	420
gaaggganga	taantgggat	ctaccaattg	attctggcaa	aacnatntct	aagattnttn	480
tgctttatgt	ggganacana	tctantctc	atttntgtct	gnanatnaca	ccctactcgt	540
gntcgancnc	gtcttcgatt	ttcgganaca	cnccantnaa	tactggcggt	ctgttggtta	600
aaaaaaaaaa	aaaa					614

<210> 36

<211> 686

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (686)

<223> n = A,T,C or G

<400> 36

gtggctggcc	cggttctcgc	cttctcccca	tcccctactt	tcctccctcc	ctccctttcc	60
ctccctcgtc	gactgttgct	tgctggtcgc	agactccctg	accctccctc	caccctcccc	120
taacctcggg	gccaccggat	tgcccttctt	ttcctgttgc	ccagcccagc	cctagtgtca	180
gggcgggggc	ctggagcagc	cggaggcact	gcagcagaag	ananaaaaga	cacgacnaac	240
ctcagctcgc	cagtcgggtc	gctngcttcc	cgccgcatgg	caatnagaca	gacgccgctc	300
acctgctctg	ggcacacgcg	accogtgggt	gatttggcct	tcagtggcat	cacccttatg	360
ggtatttctt	aatcagcgct	tgcaaagatg	gttaacctat	gctacgccag	ggagatacag	420
gagactggat	tggaacattt	ttgggggtcta	aaggctctgt	tgggggtgcaa	cactgaataa	480
ggatgccacc	aaagcagcta	cagcagctgc	agatttcaca	gcccagtggt	gggatgctgt	540
ctcagganat	naattgataa	cctggctcat	aacacattgt	caagaatgtg	gatttcccca	600
ggatattatt	atttgtttac	cggggganag	gataactgtt	tcnctntatt	taattgaaca	660
aactnaaaca	aanctaagg	aatcc				686

<210> 37

<211> 681

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (681)

<223> n = A,T,C or G

<400> 37

gagacanaacn	naacgtcang	agaanaaaaag	angcatggaa	cacaanccag	gcncgatggc	60
caccttccca	ccagcancca	gcgcccccca	gcngccccca	ngnccggang	accangactc	120
cancctgnat	caatctganc	tctattctctg	gccccatncc	acctcgagg	tggangccgn	180
aaaggtcgca	cmnncagaga	agctgctgcc	ancaccancc	gccccnnccc	tgncgggctn	240
nataggaac	tggtgaccnn	gctgcanaat	tcatacagga	gcacgcgang	ggcannnct	300
cacactgagt	tnnngatgan	gcctnaccan	ggacctnccc	cagcnnattg	annacnggac	360
tgccggaggaa	ggaagacccc	gnacnggatc	ctggccggcn	tgccaccccc	ccacccttag	420
gattatnccc	cttgactgag	tctctgaggg	gctacccgaa	ccgcctcca	ttccctacca	480
natnntgctc	natcgggact	gacangctgg	ggatnggagg	ggctateccc	cancatcccc	540

tnanaccaac	agcnacngan	natnggggct	ccccngggtc	ggngcaacnc	tectncaccc	600
cggcgcnggc	cttcgggtgt	gtcctccntc	aacnaattcc	naaanggcgg	gccccccngt	660
ggactcctcn	ttgttccctc	c				681

<210> 38

<211> 687

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(687)

<223> n = A,T,C or G

<400> 38

canaaaaaaaa	aaaacatggc	cgaaaccagn	aagctgcgcg	atggcgccac	ggccccctctt	60
ctccccggcct	gtgtccggaa	ggtttccctc	cgaggcgccc	cggtcccgcc	aagcggagga	120
gagggcgggga	cntgcggggg	cggagctca	naggccctgg	ggcgcgtctg	ctctcccgcc	180
atcgcaaggg	cggcgctaac	ctnaggcctc	cccgcgaagg	tcccnangc	ggnggcggcg	240
gggggctgtg	anaaccgcaa	aaanaacgct	gggcgcgcng	cgaaaccgct	cacccccgcg	300
aaggananac	ttccacagan	gcagcgtttc	cacagcccan	agccacnttt	ctagggtgat	360
gcacccccagt	aagtccctgn	cggggaagct	caccgctgtc	aaaaaanctc	ttcgctccac	420
cggcgccacna	aggggangan	ggcanganc	tgccgcccgc	acaggtcatc	tgatcacgtc	480
gcccccccta	ntctgctttt	gtgaatctcc	actttgttca	accccacccg	ccgttctctc	540
ctccttgccg	cttctctna	ccttaanaac	cagcttctc	taccnatng	tanttntctc	600
gcncnngtng	aaattaatc	ggtccncgg	aacctcttnc	ctgtggcaac	tgctnaaaga	660
aactgctgtt	ctgnttactg	cngtccc				687

<210> 39

<211> 695

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(695)

<223> n = A,T,C or G

<400> 39

actagtctgg	cctacaatag	tgtgattcat	gtaggacttc	tttcatcaat	tcaaaacccc	60
tagaaaaacg	tatacagatt	atataagtag	ggataagatt	tctaacattt	ctgggctctc	120
tgacccctgc	gctagactgt	ggaaagggag	tattattata	gtatacaaca	ctgctgttgc	180
cttattagtt	ataacatgat	aggtgctgaa	ttgtgattca	caatttaaaa	acactgtaat	240
ccaaactttt	ttttttaact	gtagatcatg	catgtgaatg	ttaatgttaa	tttgttcaan	300
gttggttatgg	gtagaaaaaa	ccacatgcct	taaaatttta	aaaagcaggg	cccaaactta	360
ttagtttaaa	attaggggta	tgtttccagt	ttgttattaa	ntgggtatag	ctctgtttag	420
aanaaatcna	ngaacangat	ttngaaantt	aagntgacat	tatttnccag	tgacttggtta	480
atltgaaatc	anacacggca	ccttccgttt	tggttctatt	ggnttttgaa	tccaancngg	540
ntccaaatct	tnttggaac	ngtcnnttta	acttttttac	nanatcttat	ttttttatct	600
tggaaatggc	ctatttaang	ttaaaagggg	ggggnnccac	naccattcnt	gaataaaact	660
naatatatat	ccttgggtccc	ccaaaattta	aggng			695

<210> 40

<211> 674

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)... (674)

<223> n = A,T,C or G

<400> 40

actagtagtc agttgggagt ggttgctata ccttgacttc atttatatga atttcactt	60
tattaaataa tagaaaagaa aatcccggtg cttgcagtag agttatagga cattctatgc	120
ttacagaaaa tatagccatg attgaaatca aatagtaaag gctgttctgg ctttttatct	180
tcttagctca tcttaaataa gtagtacact tgggatgcag tgcgtctgaa gtgctaata	240
gttgtaacaa tagcacaaat cgaacttagg atgtgtttct tctcttctgt gtttcgattt	300
tgatcaattc tttaattttg ggaacctata atacagtttt cctattcttg gagataaaaa	360
ttaaatggat cactgatatt taagtcattc tgcttctcat ctnaatattc catattctgt	420
attagganaa antacctccc agcacagccc cctctcaaac cccacccaaa accaagcatt	480
tggaatgagt ctcttttatt tccgaantgt ggatgggtata acccatatcn ctccaatttc	540
tgnttgggtt gggatattaat ttgaactgtg catgaaaagn ggnaatcttt nctttgggtc	600
aaantttncg ggttaatttg nctngncaaa tccaatttnc tttaagggtg tctttataaa	660
atttgcattt cngg	674

<210> 41

<211> 657

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)... (657)

<223> n = A,T,C or G

<400> 41

gaaacatgca agtaccacac actgtttgaa ttttgacaaa aaagtgactg tagggatcag	60
gtgatagccc cggaatgtac agtgtcttgg tgcaccaaga tgccttctaa aggetgacat	120
accttggggc cctaattggg cagagagtat agccctagcc cagtgggtgac atgaccactc	180
cctttgggag gctgaagtta aagggaatgg tatgtgtttt ctcattggaag cagcacatga	240
atnggtnaca ngatgttaaa ntaagntct antttgggtg tcttgtcatt tgaaaaantg	300
acacactcct ancanctggg aaaggggtgc tggaagccat ggaagaactc taaaaacatt	360
agcatgggct gatctgatta ctctctggca tcccgctcac ttttatggga agtcttatta	420
naaggatggg ananttttcc atatccttgc tgttggaact ctggaacact ctctaaattt	480
ccctctatta aaaatcactg nccttactac acttctcctc tgaanggaata gaaatggacc	540
tttctctgac ttagttcttg gcatggganc cagcccaaat taaaatctga cttntccggt	600
ttctccngaa ctcacctact tgaattggta aaacctcctt tggaattagn aaaaacc	657

<210> 42

<211> 389

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)... (389)

<223> n = A,T,C or G

<400> 42

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actagtgtctg aggaatgtaa acaagtttgc tgggccttgc gagacttcac caggttggtt      60
cgatagctca cactcctgca ctgtgcctgt caccaggaa tgtctttttt aattagaaga      120
caggaagaaa acaaaaacca gactgtgtcc cacaatcaga aacctccgtt gtggcagang      180
ggccttcacc gccaccaggg tgtcccgcc gacagggaga gactccagcc ttctgaggcc      240
atcctgaaga attcctgttt gggggttgtg aaggaaaatc acccggtttt aaaaagatgc      300
tggtgcctgc ccgcgtngtn gggaaggac tggtttcctg gtgaatttct taaaagaaaa      360
atattttaag ttaagaaaa aaaaaaaa      389

```

<210> 43

<211> 279

<212> DNA

<213> Homo sapien

<400> 43

```

actagtgaca agtccttggc cttgagatgt cttctcgtta aggagatggg ccttttgagg      60
gtaaaggata aatgaatga gttctgtcat gattcactat tctagaactt gcatgacctt      120
tactgtgta gctctttgaa tgttcttgaa attttagact ttctttgtta acaataata      180
tgctcttacc attgtataaa agctgttatg tgcaacagtg tggagatcct tgtctgattt      240
aataaaatac ttaaacactg aaaaaaaaa aaaaaaaaa      279

```

<210> 44

<211> 449

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(449)

<223> n = A,T,C or G

<400> 44

```

actagtagca tcttttctac aacgttaaaa ttgcagaagt agcttatcat taaaaaacia      60
caacaacaac aataacaata aatcctaagt gttaatcagt tattctaccc cctaccaagg      120
atatcagcct gttttttccc ttttttctcc tgggaataat tgtgggcttc ttcccaaatt      180
tctacagcct ctttctctt ctcattgctg agcttccctg tttgcacgca tgcgttgtgc      240
aagantgggc tgtttngctt ggantnccgt ccnagtggaa ncatgcttcc cctgtgtact      300
gttgggaagaa actcaaacct tcnancccta ggtgttncca ttttgtcaag tcatcactgt      360
atttttgtac tggcattaac aaaaaaagaa atnaaatatt gttccattaa actttaataa      420
aactttaaaa gggaaaaaaa aaaaaaaaa      449

```

<210> 45

<211> 559

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(559)

<223> n = A,T,C or G

<400> 45

```

actagtgtgg gggaatcacg gacacttaaa gtcaatctgc gaaataattc ttttattaca      60
cactcactga agtttttgag tcccagagag ccattctatg tcaaacattc caagtactct      120
ttgagagccc agcattacat caacatgcc gtgcagtcca aaccgaagtc cgcaggcaaa      180
tttgaagctt tgcttgtcat tcaaacagat gaaggcaaga gtattgctat toactaatt      240

```

ggtgaagctc	ttggaaaaaa	ttnactagaa	tactttttgt	gttaagttaa	ttacataagt	300
tgtattttgt	taactttatc	tttctacact	acaattatgc	ttttgtatat	atattttgta	360
tgatggatat	ctataattgt	agattttggt	tttacaagct	aatactgaag	actcgactga	420
aatattatgt	atctagccca	tagtattgta	cttaactttt	acagggtgaa	aaaaaaattc	480
tgtgtttgca	ttgattatga	tattctgaat	aaatatggga	atatatttta	atgtgggtaa	540
aaaaaaaaaa	aaaaaggaa					559

<210> 46
 <211> 731
 <212> DNA
 <213> Homo sapien

 <220>
 <221> misc_feature
 <222> (1)... (731)
 <223> n = A,T,C or G

<400> 46						
actagttcta	gtaccatggc	tgatcatagat	gcaaccatta	tattccattt	agtttcttcc	60
tcaggttccc	taacaattgt	ttgaaactga	atatatatgt	ttatgtatgt	gtgtgtgttc	120
actgtcatgt	atatggtgta	tatgggatgt	gtgcagtgtt	cagttatata	tatattcata	180
tatacatatg	catatatatg	tataatatac	atatatacat	gcatacactt	gtataatata	240
catatatata	cacatatatg	cacacatatn	atcactgagt	tccaaagtga	gtcttttatt	300
ggggcaattg	tattctctcc	ctctgtctgc	tcactgggcc	tttgcaagac	atagcaattg	360
cttgatttcc	tttgataag	agtcttatct	tcggcactct	tgactctagc	cttaacttta	420
gatttctatt	ccagaatacc	tctcatatct	atcttaaaac	ctaaganggg	taaagangtc	480
ataagattgt	agtatgaaag	antttgctta	gttaaattat	atctcaggaa	actcattcat	540
ctacaaatta	aattgtaaaa	tgatggtttg	ttgtatctga	aaaaatgttt	agaacaagaa	600
atgtaactgg	gtacctgtta	tatcaaagaa	cctcnattta	ttaagtctcc	tcatagccan	660
atccttatat	ngccctctct	gacctgantt	aatananact	tgaataatga	atagttaatt	720
taggnttggg	c					731

<210> 47
 <211> 640
 <212> DNA
 <213> Homo sapien

 <220>
 <221> misc_feature
 <222> (1)... (640)
 <223> n = A,T,C or G

<400> 47						
tgcgngccgg	tttggccctt	ctttgtanga	cactttcate	cgccctgaaa	tcttcccgat	60
cgtttaataac	tcctcaggtc	cctgcctgca	cagggttttt	tcttantttg	ttgcctaaca	120
gtacacaaaa	tgtgacatcc	tttccaat	atngattnct	tcataaccaca	tcntcnatgg	180
anacgactnc	aacaattttt	tgatnaccen	aaanactggg	ggetnnaana	agtacantct	240
ggagcagcat	ggacctgtcn	gcnactaang	gaacaanagt	nntgaacatt	tacacaacct	300
ttggtatgtc	ttactgaaag	anagaaacat	gcttctnncc	ctagaccacg	aggncaaccg	360
caganattgc	caatgccaaag	tccgagcggt	tagatcagggt	aatacattcc	atggatgcac	420
tacatacntt	gtccccgaaa	nanaagatgc	cctaanggct	tcttcanact	ggtccngaaa	480
acancatcac	ctggtgcttg	ganaacanac	tctttggaag	atcatctggc	acaagttccc	540
cccagtgggg	tttnccttgg	cacctanctt	accanatcna	ttcggaancc	attctttgcc	600
ntggcnttnt	nttgggacca	ntcttctcac	aactgnaccc			640

<210> 48
 <211> 257
 <212> DNA
 <213> Homo sapien

<400> 48

actagtatat gaaaatgtaa atatcacttg tgtactcaaa caaaagttag tcttaagctt	60
ccaccttgag cagccttgga aacctaacct gcctctttta gcataatcac attttctaaa	120
tgattttctt tgttcctgaa aaagtgtttt gtattagtgt tacatttggt ttttggaaga	180
ttatatttgt atatgtatca tcataaaaata tttaaataaa aagtatcttt agagtgaata	240
aaaaaaaaa aaaaaaa	257

<210> 49
 <211> 652
 <212> DNA
 <213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (652)

<223> n = A,T,C or G

<400> 49

actagttcag atgagtggct gctgaagggg ccccttgct attttcatta taaccaatt	60
tcacttatt tgaactctta agtcataaat gtataatgac ttatgaatta gcacagtaa	120
gttgacacta gaaactgccc atttctgtat tacactatca aataggaaac attggaaga	180
tggggaaaaa aatcttattt taaaatggct tagaaagttt tcagattact ttgaaaattc	240
taaacttctt tctgtttcca aaacttgaaa atatgtagat ggactcatgc attaagactg	300
ttttcaaagc tttctcaca tttttaaagt gtgattttcc ttttaataa catatttatt	360
ttctttaaag cagctatata ccaacccatg actttggaga tatacctatn aaaccaatat	420
aacagcangg ttattgaagc agctttctca aatgttgctt cagatgtgca agttgcaaat	480
tttattgtat ttgtanaata caattttgt tttaaactgt atttcaatct atttctcaa	540
gatgcttttc atatagagtg aaatatccca ngataactgc ttctgtgtcg tcgcatattga	600
cgcataactg cacaatgaa cagtgtatag ctcttggttg tgcattnacc cc	652

<210> 50
 <211> 650
 <212> DNA
 <213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (650)

<223> n = A,T,C or G

<400> 50

ttgcgctttg attttttttag ggcttggtgc ctgtttcact tatagggtct agaagcttg	60
tggttagtaa aaaggagatg cccaatatc aaagctgcta aatgttctct ttgccataaa	120
gactccgtgt aactgtgtga acacttgga tttttctct ctgtcccgag gtctgcgtct	180
gctttctttt ttgggttctt tctagaagat tgagaaatgc atatgacagg ctgagancac	240
ctcccaaac acacaagctc tcagccacan gcagcttctc cacagcccca gcttcgcaca	300
ggctcctgga nggctgctg ggggaggcag acatgggagt gccaagggtg ccagatggtt	360
ccaggactac aatgtcttta tttttaactg tttgccactg ctgccctcac ccctgcccgg	420
ctctggagta cgtctgccc canacaagtg ggantgaaat ggggggtggg gggaaactg	480
attcccantt aggggggtgc taactgaaca gtagggatan aaggtgtgaa cctgngaant	540

gcttttataa attatnttcc ttgttanatt tatttttttaa tttaattctt gttnaactgc 600
ccngggaaaa ggggaaaaaa aaaaaaaat tctntttaaa cacatgaaca 650

<210> 51
<211> 545
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)...(545)
<223> n = A,T,C or G

<400> 51
tggcgtgcaa ccagggtagc tgaagtttgg gtctgggact ggagattggc cattaggcct 60
cctganattc cagctccctt ccaccaagcc cagtcttgct acgtggcaca gggcaaacct 120
gactcccttt gggectcagt tccccctccc ctccatgana tgaaaagaat actacttttt 180
cttggtggtc taacnttget ggacncaaag tgtngtcatt attgttgat tgggtgatgt 240
gtncaaaact gcagaagctc actgcctatg agaggaanta agagagatag tggatganag 300
ggacanaagg agtcattatt tggatatagat ccaccntcc caacctttct ctcctcagtc 360
cctgcncctc atgtntctgg tntggtagt cctttgtgcc accanccatc atgctttgca 420
ttgctgccat cctgggaagg ggggtgnatcg tctcacaact tgttgatcgc gtttganatg 480
catgctttct tnatnaaaca aanaaannaa tgtttgacag ngtttaaaat aaaaaanaaa 540
caaaa 545

<210> 52
<211> 678
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)...(678)
<223> n = A,T,C or G

<400> 52
actagtagaa gaactttgcc gcttttgtgc ctctcacagg cgcctaaagt cattgccatg 60
ggaggaagac gatttggggg gggagggggg gggggcangg tccgtggggc ttccctant 120
ntatctccat ntccantggn cnntgtgcgc tctccctcg tcnattnga anttantccc 180
tggneccenn nccctctcnn nccnncnct cccccctcg ncnctccnn cttttntan 240
ncttcccat ctcnctccc cctnanngtc ccaacnccgn cagcaatnnc ncaattnctc 300
nctcncncc tcnnccggtt cttctnttct cnacntntnc ncnntnccn tgcnnntnaa 360
annctctccc cnetgcaanc gattctctcc ctcnncnnan ctntccactc cntncttctc 420
nncgctcct nttctcnnc ccacctctcn ccttcgnccc cantacnctc ncncccttn 480
cgnntcnttn nnntcctcnn accnccncc tcccttcncc cctcttctcc ceggtntntc 540
tctctccnnc nncnncnct cncnccntcc nngcgnccnt ttcgccccn cncnccntt 600
ccttctcnc cantccatcn cntntnccat nctnccncc nctcacnccc gctnccccn 660
ntctctttca caengtec 678

<210> 53
<211> 502
<212> DNA
<213> Homo sapien

<220>

<221> misc_feature
 <222> (1) ... (502)
 <223> n = A,T,C or G

<400> 53
 tgaagatcct ggtgtcgcca tgggcgcgcg ccccgcccgt tgttaccggt attgtaagaa 60
 caagcgcgtac ccaaagtctc gcttctgccg aggtgtccct gatgccaaaa ttgcattttt 120
 tgacctgggg cggaaaaang caaaantgga tgagtctccg ctttgtggcc acatgggtgc 180
 agatcaatat gagcagctgt cctctgaagc cctgnangct gcccgaaattt gtgccaataa 240
 gtacatggta aaaagtngtg gcnaagatgc ttccatatcc ggggtcggnt ccaccccttc 300
 cagctcatcc gcatcaacaa gatgttgctc tgtgctgggg ctgacaggct cccaacaggc 360
 atgcaagtgc cctttgaaa acccanggca ctgtggccag gggtcacatt gggccaattn 420
 atcatgttca tccgcaccaa ctgcagaaca angaacntgt naattnaagc cctgccagg 480
 gncaanttca aatttcccg cc 502

<210> 54
 <211> 494
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (494)
 <223> n = A,T,C or G

<400> 54
 actagtccaa gaaaaatatg cttaatgtat attacaaagg ctttgtatat gttaacctgt 60
 tttaatgccaa aaagtgtgct ttgtccacaa ttctcttaag acctcttcag aaagggattt 120
 gtttgccctta atgaatactg ttgggaaaaa acacagtata atgagtgaag agggcagaag 180
 caagaaattt ctacatctta gcgactccaa gaagaatgag tatccacatt tagatggcac 240
 attatgagga ctttaatctt tccctaaaca caataatgtt ttcttttttc ttttattcac 300
 atgatttcta agtatatttt tcatgcagga cagtttttca accttgatgt acagtgactg 360
 tggttaaattt ttctttcagt ggcaacctct ataacttta aaatatgggt agcatcttgt 420
 ctgttttgaa ngggatatga cnatnaatct atcagatggg aaatcctgtt tccaagttag 480
 aaaaaaaaaa aaaa 494

<210> 55
 <211> 606
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (606)
 <223> n = A,T,C or G

<400> 55
 actagtaaaa agcagcattg ccaataatc cctaattttc cactaaaaat ataatgaat 60
 gatgttaagc tttttgaaaa gtttaggtta aacctactgt tgtagatta atgtatttgt 120
 tgcttccctt tatctggaat gtggcattag cttttttatt ttaacctctt ttaattctta 180
 ttcaattcca tgacttaagg ttggagagct aaacactggg atttttggat aacagactga 240
 cagttttgca taattataat cggcattgta catagaaagg atatggctac cttttgttaa 300
 atctgcactt tctaaatc aaaaaaggga aatgaagtat aaatcaattt ttgtataatc 360
 tgtttgaaac atgantttta tttgcttaat attangcctt tgcccttttc tgttagcttc 420
 ttgggatcct gtgtaaaact gttctcatta aacaccaaag agttaagtcc attctctggt 480

```

actagctaca aattccgttt catattctac ntaacaatth aaattaactg aaatatttct 540
anatgggtcta cttctgtcnt ataaaaacna aacttgantt nccaaaaaaa aaaaaaaaaa 600
aaaaaa 606

```

<210> 56

<211> 183

<212> DNA

<213> Homo sapien

<400> 56

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actagtatat ttaaacttac aggcttattt gtaatgtaaa ccaccatttt aatgtactgt 60
aattaacatg gttataatac gtacaatcct tccctcatcc catcacacaa ctttttttgt 120
gtgtgataaa ctgatttttg tttgcaataa aaccttgaaa aataaaaaaa aaaaaaaaaa 180
aaa 183

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<210> 57

<211> 622

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (622)

<223> n = A,T,C or G

<400> 57

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actagtcact actgtcttct ccttgtagct aatcaatcaa tattcttccc ttgcctgtgg 60
gcagtggaga gtgctgctgg gtgtacgctg cacctgccca ctgagttggg gaaagaggat 120
aatcagtgag cactgttctg ctcagagctc ctgatctacc ccaccccta ggatccagga 180
ctgggtcaaa gctgcatgaa accaggccct ggcagcaacc tgggaatggc tggaggtggg 240
agagaacctg acttctcttt cctctccct cctccaacat tactggaact ctatcctgtt 300
agggatcttc tgagcttggt tccctgctgg gtgggacaga agacaaagga gaagggangg 360
tctacaanaa gcagcccttc tttgtcctct ggggttaatg agcttgacct ananttcatt 420
gaganaccan aagcctctga ttttaattt ccntnaaatg tttgaagtnt atatntacat 480
atatatattt ctttnaatnt ttgagtcctt gatatgtctt aaaatccant ccctctgccc 540
gaaacctgaa ttaaaacct gaanaaaaat gtttncctta aagatgttan taattaattg 600
aaacttgaaa aaaaaaaaaa aa 622

```

<210> 58

<211> 433

<212> DNA

<213> Homo sapien

<400> 58

```

gaacaaattc tgattggtta tgtaccgtca aaagacttga agaaatttca tgattttgca 60
gtgtggaagc gttgaaaatt gaaagtact gcttttccac ttgctcatat agtaaaggga 120
tcctttcagc tgccagtgtt gaataatgta tcatccagag tgatgttatc tgtgacagtc 180
accagcttta agctgaacca ttttatgaat accaaataaa tagacctctt gtactgaaaa 240
catatttggt actttaatcg tgctgcttg atagaaatat ttttactggg tcttctgaat 300
tgacagtaaa cctgtccatt atgaatggcc tactgttcta ttatttggtt tgacttgaat 360
ttatccacca aagacttcat ttgtgtatca tcaataaagt tgtatgttcc aactgaaaaa 420
aaaaaaaaa aaa 433

```

<210> 59

<211> 649

<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)...(649)
<223> n = A,T,C or G

<400> 59

actagttatt atctgacttt cngggtataa tcattctaata	gagtggtgaag tagcctctgg	60
tgtcatttgg atttgcatth ctctgatgag tgatgctatc	aagcaccttt gctgggtgctg	120
ttggccatat gtgtatgttc cctggagaag tgtctgtgct	gagccttggc ccacttttta	180
attaggcgtn tgtcttttta ttactgagtt gtaaganttc	tttatatatt ctggattcta	240
gacccttate agatacatgg ttgcaaata tttctccca	ttctgtgggt tgtgttttca	300
ctttatcgat aatgtcctta gacatataat aaatttgat	tttaaaagtg acttgatttg	360
ggctgtgcaa ggtgggtca ogcttgtaat cccagcactt	tgggagactg aggtgggtgg	420
atcatatgan gangctagga gtteggaggtc agcctggcca	gcatagcgaa aacttgtctc	480
tacnaaaaat acaaaaatta gtcaggcatg gtgggtgcag	tctgtaatac cagcttctca	540
ggangctgan gcacaaggat cacttgaacc ccagaangaa	gangttgcag tganctgaag	600
atcatgccag ggcaacaaaa atgagaactt gtttaaaaaa	aaaaaaaaa	649

<210> 60
<211> 423
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)...(423)
<223> n = A,T,C or G

<400> 60

actagttcag gccttcag tcaactgacaa acatggggaa	gtgtgcccag ctggctggaa	60
acctggcagt gataccatca agcctgatgt ccaaaagagc	aaagaatatt tctccaagca	120
gaagtgagcg ctgggctgtt ttagtgccag gctgcggtgg	gcagccatga gaacaaaacc	180
tcttctgtat ttttttttc cattagtana acacaagact	cngattcagc cgaattgtgg	240
tgtcttaciaa ggcagggtt tctacaggg ggtgganaaa	acagcctttc ttcctttggt	300
aggaatggcc tgagttggcg ttgtgggcag gctactggtt	tgtatgatgt attagtagag	360
caaccatta atcttttgta gtttgatna aacttganct	gagaccttaa acaaaaaaaa	420
aaa		423

<210> 61
<211> 423
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)...(423)
<223> n = A,T,C or G

<400> 61

cgggactgga atgtaaagtg aagttcggag ctctgagcac	gggctcttcc cgccgggtcc	60
tccctcccca gacccagag ggagaggccc accccgcca	gccccgccc agccctgct	120
caggctctgag tatggctggg agtcgggggc cacaggcctc	tagctgtgct gctcaagaag	180


```

actggatcag ggtanctaca agtggccggg ccttgccctt gggattctac cctgttccta 240
atttggtggt ggggtgctgg gtccttgccc cccttttcca cactnccctc ctccngacag 300
caacctccct tggggcaatt gggcctggnt ctcnccccgn tgggtgcnacc ctttggtggt 360
ttaaggncct taaaaatggt annttttccc ntgcncgggt taaaaaagga aaaaactnaa 420
aaa 423

```

```

<210> 62
<211> 683
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)...(683)
<223> n = A,T,C or G

```

```

<400> 62
gctggagagg ggtacggact ttcttggagt tgtcccaggt tggaaatgaga ctgaactcaa 60
gaagagaccc taagagactg gggaaatggtt cctgccttca ggaaagtga agacgcttag 120
gctgtcaaca cttaaaggaa gtccccttga agcccagagt ggacagacta gacccattga 180
tggggccact ggccatggc cgtggacaag acattccngt gggccatggc acaccggggg 240
ggatcaaaat gtgtacttgt ggggtctcgc cccttgccaa aaccaaacca ntccactcc 300
tgtcnttggga ctttcttccc attccctcct ccccaaattgc acttcccctc ctccctctgc 360
ccctcctgtg tttttggaat tctgtttccc tcaaaattgt taatttttta nttttngacc 420
atgaacttat gtttggggtc nangttcccc ttaccaatgc atactaatat attaatggtt 480
atttattttt gaaatatttt ttaatgaact tggaaaaaat tnntggaatt tccttncttc 540
cntttntttt ggggggggtg gggggntggg ttaaaatttt tttggaancc cnatnggaaa 600
ttnttacttg gggccccctt naaaaaantn anttccaatt cttnnatngc cctnttccn 660
ctaaaaaaa ananannaaa aan 683

```

```

<210> 63
<211> 731
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)...(731)
<223> n = A,T,C or G

```

```

<400> 63
actagtcata aagggtgtgc gcgtcttcga cgtggcggtc ttggcgccac tgctgcgaga 60
cccggccctg gacctcaagg tcatccactt ggtgcgtgat cccgcgcggg tggcgagttc 120
acggatccgc tcgcgccacg gcctcatccg tgagagccta caggtggtgc gcagccgaga 180
ccgcgagctc accgcatgcc cttcttggag gccgcgggcc acaagcttgg cgcctnaaaa 240
gaaggcgtng ggggcccgcg aantaccacg ctctggcggc tatggaangt cctcttgcaa 300
taatattggt tnaaaanctg canaanagcc cctgcanccc cctgaactgg gntgcagggc 360
cncttacctn gtttgntgc ggttacaaag aacctgtttn ggaaaaccct nccnaaaacc 420
ttccgggaaa attntncaaa ttttnttggg ggaattnttg ggtaaacccc ccnaaaatgg 480
gaaacntttt tgcctnmaa antaaacat tnggttccgg gggccccccc ncaaaaccct 540
ttttntttt tttntgcccc cantnncccc ccggggcccc ttttttngg ggaaaanccc 600
ccccctncc nanantttta aaagggnggg anaatttttn ntncccccc gggncccccn 660
gngngtaaaa nggtttcncc ccccgaggg gnggggnnnc ctcnnaaacc cntntcnnaa 720
ccncttttn n 731

```

<210> 64
 <211> 313
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(313)
 <223> n = A,T,C or G

<400> 64
 actagttgtg caaaccacga ctgaagaaag acgaaaagtg ggaaataact tgcaacgtct 60
 gttagagatg gttgctacac atgttggtgc tgtagagaaa catcttgagg agcagattgc 120
 taaagttgat agagaatatg aagaatgcat gtcagaagat ctctcggaaa atattaaaga 180
 gattagagat aagtatgaga agaaagctac tctaattaag tcttctgaag aatgaagatn 240
 aaatgttgat catgtatata tatccatagt gaataaaatt gtctcagtaa agttgtaaaa 300
 aaaaaaaaaa aaa 313

<210> 65
 <211> 420
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(420)
 <223> n = A,T,C or G

<400> 65
 actagttccc tggcaggcaa gggcttccaa ctgaggcagt gcatgtgtgg cagagagagg 60
 caggaagctg gcagtggcag cttctgtgtc tagggagggg tgtggctccc tcttccctg 120
 tctgggaggt tggagggaag aatctaggcc ttagcttgcc ctctgccac cttccctt 180
 gtagatactg ccttaacact cctcctctc tcagctgtgg ctgccacca agccaggtt 240
 ctccgtgctc actaatttat ttccaggaaa ggtgtgtgga agacatgagc cgtgtataat 300
 atttgtttta acattttcat tgcaagtatt gaccatcatc cttggttgtg tatcgttgta 360
 acacaaatta atgatattaa aaagcatcca aacaaagccn annnnnaana nnannngaaa 420

<210> 66
 <211> 676
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(676)
 <223> n = A,T,C or G

<400> 66
 actagtttcc tatgatcatt aaactcattc tcagggttaa gaaaggaatg taaatttctg 60
 cctcaatttg tacttcatca ataagttttt gaagagtgc gatttttagt caggtcttaa 120
 aaataaaactc acaaatctgg atgcatttct aaattctgca aatgtttcct ggggtgactt 180
 aacaaggaat aatcccacaa tatacctagc tacctaatac atggagctgg ggctcaaccc 240
 actgttttta aggatttgcg cttacttgtg gctgaggaaa aataagtagt tccgagggaa 300
 gtagttttta aatgtgagct tatagatngg aaacagaata tcaacttaat tatggaaatt 360
 gtagaaaacc tgttctcttg ttatctgaat cttgattgca attactattg tactggatag 420

actccagccc attgcaaagt ctcagatata ttanctgtgt agttgaattc cttggaaatt	480
ctttttaaga aaaaattgga gtttnaaaga aataaacccc tttgttaaata gaagcttggc	540
tttttgggtga aaaanaatca tccgcaggg cttattgttt aaaaanggaa ttttaagcct	600
ccctggaaaa anttgtaata taaatgggga aaatgntggg naaaaattat ccgttagggg	660
ttaaagggaa aactta	676

<210> 67
 <211> 620
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(620)
 <223> n = A,T,C or G

<400> 67	
caccattaaa gctgcttacc aagaacttcc ccagcatttt gacttccttg tttgatagct	60
gaattgtgag caggtgatag aagagccttt ctagttgaac atacagataa tttgctgaat	120
acattccatt taatgaaggg gttacatctg ttacgaagct actaagaagg agcaagagca	180
taggggaaaa aaatctgatc agaacgcata aaactcacat gtgccccctc tactacaaac	240
agattgtagt gctgtggtgg tttattccgt tgtgcagaac ttgcaagctg agtcactaaa	300
cccaaagaga ggaaattata ggtagttaa acattgtaat ccaggaact aagtttaatt	360
cacttttgaa gtgttttgtt tttattttt ggttgtctg atttactttg ggggaaaang	420
ctaaaaaaaa agggatatca atctctaatt cagtgccac taaaagttgt ccctaaaaag	480
tctttactgg aanttattggg actttttaag ctccaggtnt tttggtcctc caaattaacc	540
ttgcatgggc cccttaaaat tgttgaangg cattcctgcc tctaagtttg gggaaaattc	600
ccccnttttn aaaatttga	620

<210> 68
 <211> 551
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(551)
 <223> n = A,T,C or G

<400> 68	
actagtagct ggtacataat cactgaggag ctatttctta acatgctttt atagaccatg	60
ctaagtctag accagtattt aagggtctaat ctacacctc cttagctgta agagtctggc	120
ttagaacaga cctctctgtg caataacttg tggccactgg aaatccctgg gccggcattt	180
gtattggggg tgcaatgact cccaagggcc aaaagagtta aaggcacgac tgggatttct	240
tctgagactg tgggtgaaact ccttccaagg ctgagggggg cagtangtgc tctgggaggg	300
actcggcacc actttgatat tcaacaagcc acttgaagcc caattataaa attgttattt	360
tacagctgat ggaactcaat ttgaaccttc aaaactttgt tagtttatcc tattatattg	420
ttaaacctaa ttacatttgt ctagcattgg atttgggtcc tgtngcatat gttttttcn	480
cctatgtgct cccctcccc nnatcttaat ttaaaccnca attttgcnat tcnccnnnnn	540
nannnnnnna a	551

<210> 69
 <211> 396
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)... (396)
 <223> n = A,T,C or G

<400> 69
 cagaaatgga aagcagagtt ttcattttctg tttataaaacg tctccaaaca aaaatggaaa 60
 gcagagtttt cattaaatcc ttttaccttt tttttttctt ggtaatcccc tcaaataaca 120
 gtatgtggga tattgaatgt taaagggata tttttttcta ttatttttat aattgtacaa 180
 aattaagcaa atgttaaaag ttttatatgc tttattaatg ttttcaaaag gtatnatata 240
 tgtgatacat tttttaagct tcagttgctt gtcttctggt actttctggt atgggctttt 300
 ggggagccan aaaccaatct acnatctctt tttgtttgcc aggacatgca ataaaattta 360
 aaaaataaat aaaaactatt nagaaattga aaaaaa 396

<210> 70
 <211> 536
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)... (536)
 <223> n = A,T,C or G

<400> 70
 actagtgcga aagcaaatat aaacatcgaa aaggcggtcc tcacgttagc tgaagatata 60
 cttcgaaga cccctgtaaa agagcccaac agtgaaaatg tagatatcag cagtggagga 120
 ggcgtgacag gctggaagag caaatgctgc tgagcattct cctgttccat cagttgccat 180
 ccactacccc gttttctctt cttgctgcaa aataaaccac tctgtccatt tttactcta 240
 aacagatatt tttgtttctc atcttaacta tccaagccac ctattttatt tgttctttca 300
 tctgtgactg cttgctgact ttatcataat tttcttcaaa caaaaaaatg tatagaaaaa 360
 tcatgtctgt gacttcattt ttaaatgnta cttgctcagc tcaactgcat ttcagttggt 420
 ttatagtcca gttcttatca acattnaaac ctatngcaat catttcaaat ctattctgca 480
 aattgtataa gaataaaagt tagaatttaa caattaaaaa aaaaaaaaaa aaaaaa 536

<210> 71
 <211> 865
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)... (865)
 <223> n = A,T,C or G

<400> 71
 gacaaagcgt taggagaaga anagaggcag ggaanactnc ccaggcacga tggccnctt 60
 cccaccagca accagcgccc cccaccagcc cccaggcccg gacgacgaag actccatcct 120
 ggattaatct nacctctntc gcctgnccca ttcctacctc ggaggtggag gccggaaagg 180
 tcncaccaag aganaantcgt ctgccaacac caaccgcccc agccctggcg ggcacganag 240
 gaaactggtg accaatctgc agaattctna gaggaanaag cnaggggccc cgcgctnaga 300
 cagagctgga tatgangcca gaccatggac nctacnccn ncaatncana cgggactgcy 360
 gaagatggan gaccncgac nngatcagge cngctnncca nccccccacc cctatgaatt 420
 attccogctg aangaatctc tgannggctt ccannaaagc gcctccccnc cnaacgnaan 480

tncaacatng	ggattanang	ctgggaactg	naaggggcaa	ancctnnaat	atccccagaa	540
acaantcttc	ccnaanaaac	tggggcncct	catnggtggn	accaactatt	aactaaaccg	600
cacgccaagn	aantataaaa	ggggggcccc	tccncggng	acccctttt	gtcccttaat	660
ganggttatc	cnccttgct	accatggtn	ccmmttctgt	ntgnatgtt	ccnctccct	720
ccnctatnt	cnagccgaac	tcnnatttnc	ccgggggtgc	natcnantng	tncnctttt	780
ttngttgncc	cngccctttc	cgnccgaacn	cgtttccccc	ttantaacgg	cacccgggg	840
aagggtgntt	ggccccctcc	ctccc				865

<210> 72

<211> 560

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(560)

<223> n = A,T,C or G

<400> 72

cctggacttg	tcttggttcc	agaacctgac	gacccggcga	cggcgacgtc	tcttttgact	60
aaaagacagt	gtccagtgt	ccngcctagg	agtctacggg	gaccgcctcc	cgcgcgccca	120
ccatgcccaa	cttctctggc	aactggaaaa	tcacccgac	ggaaaacttc	gangaattgc	180
tcnaantgct	gggggtgaat	gtgatgctna	ngaanattgc	tgtggctgca	gcgtccaagc	240
cagcagtggg	gatcnaacag	gaggagagaca	ctttctacat	caaaacctcc	accacgtgc	300
gcaccacaaa	gattaacttc	nnngttgggg	aggantttga	ggancaaact	gtggatngga	360
ngcctgtnaa	aacctggtga	aatgggagaa	tganaataaa	atggtctgtg	ancanaaact	420
cctgaaagga	gaaggccccc	anaactcctg	gaccngaaaa	actgaccnc	cnatngggga	480
actgatnctt	gaaccctgaa	cgggcgggat	ganccttttt	tnttgcncnc	naanggggtc	540
tttcnntttc	cccaaaaaaa					560

<210> 73

<211> 379

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(379)

<223> n = A,T,C or G

<400> 73

ctggggancc	ggcggtnngc	nccatntcnn	gncgcgaagg	tggcaataaa	aanccnctga	60
aaccgcncaa	naaacatgcc	naagatatgg	acgaggaaga	tnngctttc	nngnacaanc	120
gnanngagga	acanaacaaa	ctcnangagc	tctcaagcta	atgccgcggg	gaagggggccc	180
ttggccacnn	gtggaattaa	gaaatctggc	aaanngtann	tgttccttgt	gcctnangag	240
ataagngacc	ctttatttca	tctgtattta	aacctctctn	ttccctgnca	taacttcttt	300
tnccacgtan	agntggaant	anttggtgtc	ttggactgtt	gtncatttta	gannaaactt	360
ttgttcaaaa	aaaaaataa					379

<210> 74

<211> 437

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature
 <222> (1)...(437)
 <223> n = A,T,C or G

<400> 74

actagttcag	actgccacgc	caaccccgaga	aaatacccca	catgccagaa	aagtgaagtc	60
ctagggtgtt	ccatctatgt	ttcaatctgt	ccatctacca	ggcctcgca	taaaaacaaa	120
acaaaaaac	gctgccaggt	tttanaagca	gttctgggtc	caaaaccatc	aggatcctgc	180
caccaggggt	cttttgaaat	agtaccacat	gtaaaaggga	atttggcttt	cacttcatct	240
aatcactgaa	ttgtcaggct	ttgattgata	attgtagaaa	taagtagcct	tctgttgtgg	300
gaataagtta	taatcagtat	tcactctctt	gttttttgtc	actcttttct	ctctnattgt	360
gtcatttgta	ctgtttgaaa	aatatttctt	ctataaaatt	aaactaacct	gccttaaaaa	420
aaaaaaaaa	aaaaaaaaa					437

<210> 75
 <211> 579
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(579)
 <223> n = A,T,C or G

<400> 75

ctccgtcgcc	gccaaagatga	tgtgcggggc	gccctccgcc	acgcagccgg	ccaccgcga	60
gaccagcac	atcgccgacc	agggtgaggc	ccagcttgaa	gagaaagaaa	acaagaagtt	120
ccctgtgttt	aaggccgtgt	cattcaagag	ccagggtggc	gcggggacaa	actacttcat	180
caagggtcac	gtcggcgacg	aggacttctg	acacctgcga	gtgttccaat	ctctccctca	240
tgaaaacaag	cccttgacct	tatctaacta	ccagaccaac	aaagccaagc	atgatgagct	300
gacctatttc	tgatcctgac	tttggaacaag	gcccttcagc	cagaagactg	acaaagtcac	360
cctccgtcta	ccagagcgtg	cacttgtgat	cctaaaataa	gcttcatctc	cgggctgtgc	420
ccttgggggtg	gaaggggcan	gatctgcact	gcttttgcac	ttctcttcct	aaatttcatt	480
gtgttgattc	tttcttcca	atagggtgatc	tttattactt	tcagaatatt	ttccaaatna	540
gatatatttt	naaaatcctt	aaaaaaaaa	aaaaaaaaa			579

<210> 76
 <211> 666
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(666)
 <223> n = A,T,C or G

<400> 76

gtttatctta	tctctccaac	cagattgtca	gctccttgag	ggcaagagcc	acagtatatt	60
tccctgtttc	ttccacagt	cctaataata	ctgtggaact	aggttttaat	aatttttta	120
ttgatgttgt	tatgggcagg	atggcaacca	gaccattgtc	tcagagcagg	tgtctggctct	180
ttcctggcta	ctccatgttg	gctagcctct	ggtaacctct	tacttattat	cttcaggaca	240
ctcactacag	ggaccaggga	tgatgcaaca	tccttgtctt	tttatgacag	gatgtttgct	300
cagcttctcc	aacaataaaa	agcacgtggg	aaaacacttg	cggatattct	ggactgtttt	360
taaaaaatat	acagtttacc	gaaaatcata	ttatcttaca	atgaaaagga	ntttatagat	420
cagccagtga	acaacctttt	cccaccatac	aaaaattcct	tttcccgaa	gaaaanggct	480

ttctcaataa nectcacttt cttaanatct tacaagatag ccccganatc ttatcgaaac	540
tcatttttagg caaatatgan ttttattgtn cgttacttgt ttcaaaattt ggtattgtga	600
atatcaatta ccaccccat ctcccatgaa anaaanggga aanggtgaan ttcntaancg	660
cttaaa	666

<210> 77

<211> 396

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (396)

<223> n = A,T,C or G

<400> 77

ctgcagcccg ggggatccac taatctacca nggttatttg gcagctaatt ctanatttgg	60
atcattgccc aaagttgcac ttgctggtct cttgggattt ggccttggaa aggtatcata	120
catanganta tgccanaata aattccattt ttttgaaaat canctccttg ggcctggttt	180
tgggtccacag cataacangc actgcctcct tacctgtgag gaatgcaaaa taaagcatgg	240
attaagtgg aaggagact ctgagccttc agcttcctaa attctgtgtc tgtgactttc	300
gaagtttttt aaacctctga atttgtacac attttaaatt tcaagtgtac tttaaaataa	360
aatacttcta atgggaacaa aaaaaaaaaa aaaaaa	396

<210> 78

<211> 793

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (793)

<223> n = A,T,C or G

<400> 78

gcattcctagc cgccgactca cacaaggcag gtgggtgagg aaatccagag ttgccatgga	60
gaaaattcca gtgtcagcat tcttgtcctc tgtggccctc tcctacactc tggccagaga	120
taccacagtc aaacctggag ccaaaaagga cacaaggac tctcgacce aactgccccca	180
gaccctctcc agaggttggg gtgaccaact catctggact cagacatagt aagaagctct	240
atataaatcc aagacaagca acaaaccctt gatgattatt catcacttgg atgagtgcc	300
acacagtcna gctttaaaga aagtgtttgc tgaaaataaa gaaatccaga aattggcaga	360
gcagtttgtc ctctcaatc tggtttatga aacaactgac aaacaccttt ctctgatgg	420
ccagtatgtc ccaggattat gttgttgac ccatctctga cagttgaagc cgatatectg	480
ggaagatatt cnaaccgtct ctatgcttac aaactgcaga tacgctctgt tgcttgacac	540
atgaaaaagc tctcaagttg ctnaaaatga attgtgaaga aaaaaatctc cagccttctg	600
tctgtcggct tgaaaattga aaccagaaaa atgtgaaaaa tggctattgt ggaacanatn	660
gacacctgat taggttttgg ttatgttcac cactattttt aanaaaanan nttttaaaat	720
ttggttcaat tntctttttn aaacaatntg tttctacntt gnganctgat ttctaaaaaa	780
aataatnttt ggc	793

<210> 79

<211> 456

<212> DNA

<213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(456)
 <223> n = A,T,C or G

<400> 79

actagtatgg	ggtgggaggc	cccacccttc	tcccctagge	gctgttcttg	ctccaaaggg	60
ctccgtggag	agggactggc	agagctgang	ccacctgggg	ctggggatcc	cactcttctt	120
gcagctgttg	agcgaccta	accactgggc	atgccccac	ccctgctctc	cgcacccgct	180
tcctcccgc	cccangacca	ggctacttct	cccctcctct	tgccctccctc	ctgccccctgc	240
tgccctctgat	cgtangaatt	gangantgtc	cgccttcttg	gctganaatg	gacagtggca	300
ggggctggaa	atgggtgtgt	gtgtgtgtgt	gtgtgtgtgt	gtgtgtgtgt	gcnccccccc	360
tgcaagaccg	agattgaggg	aaancatgtc	tgctgggtgt	gaccatgttt	cctctccata	420
aantnccccct	gtgacnctca	naaaaaaaaa	aaaaaa			456

<210> 80
 <211> 284
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(284)
 <223> n = A,T,C or G

<400> 80

ctttgtacct	ctagaaaaga	taggtattgt	gtcatgaaac	ttgagtttaa	attttatata	60
taaaactaaa	agtaatgtc	acttttagca	cacatactaa	aattggaacc	atactgagaa	120
gaatagcatg	acctccgtgc	aaacaggaca	agcaaatttg	tgatgtgttg	attaaaaaga	180
aataaataaa	tgtgtatatg	tgtaacttgt	atgtttatgt	ggaatacaga	ttgggaaata	240
aatgtatttt	cttactgtga	aaaaaaaaaa	aaaaaaaaaa	aana		284

<210> 81
 <211> 671
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(671)
 <223> n = A,T,C or G

<400> 81

gccaccaaca	ttccaagcta	ccctgggtac	ctttgtgcag	tagaagctag	tgagcatgtg	60
agcaagcggg	gtgcacacgg	agactcatcg	ttataattta	ctatctgcc	agagtagaaa	120
gaaaggctgg	ggatatttgg	gttggcttgg	ttttgatttt	ttgcttgttt	gtttgttttg	180
tactaaaaca	gtattatctt	ttgaatatcg	tagggacata	agtatataca	tgttatccaa	240
tcaagatggc	tagaatggg	cctttctgag	tgtctaaaac	ttgacacccc	tggtaaatct	300
ttcaacacac	ttccactgcc	tgogtaatga	agttttgatt	catttttaac	cactggaatt	360
tttcattggc	gtcattttca	gttagatnat	tttgacattt	gagattaaaa	tgccatgtct	420
atttgattag	tcttattttt	ttatttttac	aggcttatca	gtctcactgt	tggtgtcat	480
tgtgacaaag	tcaaataaac	ccccnaggac	aacacacagt	atgggatcac	atattgtttg	540
acattaagct	ttggccaaaa	aatgttgc	gtgttttacc	tcgacttgct	aaatcaatan	600
canaaaggct	ggctnataat	gttgggtggg	aaataattaa	tnantaacca	aaaaaaaaan	660
aaaaaaaaaa	a					671

<210> 82
 <211> 217
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(217)
 <223> n = A,T,C or G

<400> 82
 ctgcagatgt ttcttgaatg ctttgtcaaa ttaanaaagt taaagtgcaa taatgtttga 60
 agacaataag tgggtgtgta tcttgtttct aataagataa acttttttgt ctttgcttta 120
 tcttattagg gagttgtatg tcagtgtata aaacatactg tgttgtataa caggcttaat 180
 aaattcttta aaaggaaaaa aaaaaaaaaa aaaaaaa 217

<210> 83
 <211> 460
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(460)
 <223> n = A,T,C or G

<400> 83
 cgcgagtggg agcaccagga tctcgggctc ggaacgagac tgcacggatt gttttaagaa 60
 aatggcagac aaaccagaca tgggggaaat cgccagcttc gatnaggcca agctgaanaa 120
 aacggagacg caggagaaga acaccctgcc gaccaaagag accattgagc angagaagcg 180
 gagtgaatt tcctaagatc ctggaggatt tcctaccccc gtctctctcg agacccagct 240
 cgtgatgtgg aggaagagcc acctgcaaga tggacacgag ccacaagctg cactgtgaac 300
 ctgggcactc cgcgccgatg ccaccggcct gtgggtctct gaagggaccc cccccaatcg 360
 gactgccaaa ttctccggtt tgccccggga tattatacaa nattatttgt atgaataatg 420
 annataaaac acacctcgtg gcancaaana aaaaaaaaaa 460

<210> 84
 <211> 323
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(323)
 <223> n = A,T,C or G

<400> 84
 tgggtgatct tggctctgtg gagctgctgg gacgggatct aaaagactat tctggaagct 60
 gtggtccaan gcattttgct ggcttaacgg gtcccggaac aaaggacacc agctctctaa 120
 aattgaagtt taccoganat aacaatcttt tgggcagaga tgcctatttt aacaaacncc 180
 gtccctgcgc aacaacnaac aatctctggg aaataccggc catgaacntg ctgtctcaat 240
 cnancatctc tctagctgac cgatcatatc gtcccagatt actacanatc ataataattg 300
 atttctctgta naaaaaaaaaa aaa 323

<210> 85
 <211> 771
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(771)
 <223> n = A,T,C or G

<400> 85
 aaactgggta ctcaacactg agcagatctg ttctttgagc taaaaacat gtgctgtacc 60
 aanagtttgc tcctggctgc tttgatgtca gtgctgtac tccacctctg cggcgaatca 120
 gaagcaagca actttgactg ctgtcttgga tacacagacc gtattcttca tcctaaattt 180
 attgtgggct tcacacggca gctggccaat gaaggctgtg acatcaatgc tatcatcttt 240
 cacacaaaga aaaagttgtc tgtgtgcgca aatccaaaac agacttgggt gaaatatatt 300
 gtgogtctcc tcagtaaaaa agtcaagaac atgtaaaaac tgtggctttt ctggaatgga 360
 attggacata gccaagaac agaaagaact tgctggggtt ggagggttca cttgcacatc 420
 atgganggtt tagtgcttat cttatttgtg cctcctggac ttgtccaatt natgaagtta 480
 atcatattgc atcatanttt gctttgttta acatcacatt naaattaaac tgtattttat 540
 gttatttata gctntaggtt ttctgtgttt aactttttat acnaantttc cttaaactatt 600
 ttggtntant gcaanttaaa aattatattt ggggggggaa taaatatggg antttctgca 660
 gccacaagct ttttttaaaa aaccantaca nccnngttaa atggtnggtc ccnaatgggt 720
 tttgcttttn antagaaaat ttnttagaac natttgaaaa aaaaaaaaaa a 771

<210> 86
 <211> 628
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(628)
 <223> n = A,T,C or G

<400> 86
 actagtttgc tttacatttt tgaaaagtat tatttttgtc caagtgttta tcaactaaac 60
 cttgtgtag gtaagaatgg aatttattaa gtgaatcagt gtgaccttc ttgtcataag 120
 attatcttaa agctgaagcc aaaatatgct tcaaaagaaa angactttat tgttcattgt 180
 agttcataca ttcaaagcat ctgaactgta gtttctatag caagccaatt acatccataa 240
 gtggagaang aaatagatta atgtcnaagt atgattgggtg gagggagcaa ggttgaagat 300
 aatctggggt tgaaattttc tagttttcat tctgtacatt tttagttna catcagattt 360
 gaaatattaa tgtttacctt tcaatgtgtg gtatcagctg gactcantaa cacccttctc 420
 ttccctnngg gatggggaat ggattattgg aaaatggaaa gaaaaaagta cttaaagcct 480
 tcctttcnca gtttctggct cctacctac tgatttancc agaataagaa aacattttat 540
 catcntctgc tttattccca ttaatnaant tttgatgaat aaatctgctt ttatgcnnac 600
 ccaaggaatt nagtggnttc ntcttgt 628

<210> 87
 <211> 518
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature

<222> (1) ... (518)

<223> n = A, T, C or G

<400> 87

tttttttttt	tttttagaga	gtagttcagc	ttttatttat	aaattttattg	cctgtttttat	60
tataacaaca	ttatactgtt	tatggtttaa	tacatatggg	tcaaaatgta	taatacatca	120
agtagtacag	ttttaaaatt	ttatgcttaa	aacaagtttt	gtgtaaaaaa	tgtagatata	180
ttttacatgg	caaatcaatt	tttaagtcac	cctaaaaatt	gatttttttt	tgaaatttaa	240
aaacacattt	aatttcaatt	tctctcttat	ataaccttta	ttactatagc	atggtttcca	300
ctacagttta	acaatgcagc	aaaattccca	tttcacggta	aattgggttt	taagcggcaa	360
ggttaaaatg	ctttgaggat	cctnaatacc	ctttgaactt	caaatgaagg	ttatggttgt	420
naatttaacc	ctcatgccat	aagcagaagc	acaagtttag	ctgcattttg	ctctaaactg	480
taaaancgag	ccccccgttg	aaaaagcaaa	agggaccc			518

<210> 88

<211> 1844

<212> DNA

<213> Homo sapien

<400> 88

gagacagtga	atcctagtat	caaaggattt	ttggcctcag	aaaaagttgt	tgattatttt	60
tattttat	tatttttcca	gactccgtct	caaaaaaaaa	aaaaaaaaaa	agaatcacaa	120
ggtagttgct	aaagcatttt	gagctgcttg	gaaaaaggga	agtagttgca	gtagagtttc	180
ttccatcttc	ttgggtgctg	gaagccatat	atgtgtcttt	tactcaagct	aaggggtata	240
agcttatgtg	ttgaatttgc	tacatctata	tttcacatat	tctcacaata	agagaatttt	300
gaaatagaaa	tatcatagaa	catttaagaa	agtttagtat	aaataatatt	ttgtgtgttt	360
taatcccttt	gaagggatct	atccaaagaa	aatattttac	actgagctcc	ttcctacacg	420
tctcagtaac	agatccctgtg	ttagtctttg	aaaatagctc	atttttttaa	tgtagtgag	480
tagatgtagc	atacatatga	tgtataatga	cgtgtattat	gttaacaatg	tctgcagatt	540
ttgtaggaat	acaaaacatg	gcctttttta	taagcaaaac	gggccaatga	ctagaataac	600
acatagggca	atctgtgaat	atgtattata	agcagcattc	cagaaaagta	gttgggtgaaa	660
taattttcaa	gtcaaaaagg	gatattgaaa	gggaattatg	agtaacctct	attttttaag	720
ccttgctttt	aaattaaacg	ctacagccat	tttaagccttg	aggataataa	agcttgagag	780
taataatgtt	aggttagcaa	aggttttagat	gtatcacttc	atgcatgcta	ccatgatagt	840
aatgcagctc	ttogagtcac	ttctgggtcat	tcaagatatt	cacccttttg	cccatagaaa	900
gcaccctacc	tcacctgctt	actgacattg	tcttagctga	tcacaagatc	attatcagcc	960
tccattattc	cttactgtat	ataaaataca	gagttttata	ttttcccttc	ttcgtttttc	1020
accatattca	aaacctaaat	ttgtttttgc	agatggaatg	caaagtaatc	aagtgttcgt	1080
gctttcacct	agaaggggtg	ggctcctgaag	gaaagagggtc	cctaaatatc	ccccaccctg	1140
gggtgctcctc	cttccctggg	accctgacta	ccagaagtca	gggtgctagag	cagctggaga	1200
agtgccagcag	cctgtgcttc	cacagatggg	gggtgctgctg	caacaaggct	ttcaatgtgc	1260
ccatcttagg	gggagaagct	agatccctgtg	cagcagcctg	gtaagtcctg	aggaggttcc	1320
attgctcttc	ctgctgctgt	cctttgcttc	tcaacggggc	tcgctctaca	gtctagagca	1380
catgcagcta	acttgtgcct	ctgcttatgc	atgaggggta	aattaacaac	cataaccttc	1440
atttgaagtt	caaaggtgta	ttcaggatcc	tcaaagcatt	tttaaccttg	cgcttaaaac	1500
ccaattttacc	gtgaaatggg	aattttgctg	cattgtttaa	ctgtagtgga	aacctgcta	1560
tagtaataaa	gggtatataa	gagagaaatt	gaaattaaat	gtgtttttta	atttcaaaaa	1620
aaaatcaatc	tttaggatga	cttaaaaatt	gatttgccat	gtaaaatgta	tctgcatttt	1680
ttacacaaaa	cttgttttta	gcataaaatt	ttaaaactgt	actacttgat	gtattatata	1740
ttttgaacca	tatgtattaa	accataaaca	gtataatgtt	gttataataa	aacaggcaat	1800
aaattttataa	ataaaagctg	aaaaaaaaaa	aaaaaaaaaa	aaaa		1844

<210> 89

<211> 523

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(523)

<223> n = A,T,C or G

<400> 89

tttttttttt	tttttttagt	caatccacat	ttattgatca	cttattatgt	accaggcact	60
gggataaaga	tgactgtag	tcactcacag	taaggaagaa	aactagcaaa	taagacgatt	120
acaatatgat	gtagaaaatg	ctaagccaga	gatatagaaa	ggtcctattg	ggtccttctg	180
tcaccttgtc	tttcacatc	cctacccttc	acaggccttc	cctccagctt	cctgcccccg	240
ctccccactg	cagatcccct	gggattttgc	ctagagctaa	acgagganat	gggccccctg	300
gccctggcat	gacttgaacc	caaccacaga	ctgggaaagg	gagcctttcg	anagtggatc	360
actttgatna	gaaaacacat	aggggaattga	agagaaantc	cccaaattggc	cacccgtgct	420
ggtgtcaag	aaaagtgtgc	agaatggata	aatgaaggat	caagggaatt	aatanaatgaa	480
taattgaatg	gtggctcaat	aagaatgact	ncnttgaatg	acc		523

<210> 90

<211> 604

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(604)

<223> n = A,T,C or G

<400> 90

ccagtgtggt	ggaatgcaaa	gattaccccg	gaagctttcg	agaagctggg	attccctgca	60
gcaaaggaaa	tagccaatat	gtgtcggttc	tatgaaatga	agccagaccg	agatgtcaat	120
ctcaccacc	aactaaatcc	caaagtcaaa	agcttcagcc	agtttatctc	agagaaccag	180
gggagccttc	aagggcatgt	agaaaatcag	ctgttcagat	aggcctctgc	accacacagc	240
ctctttcttc	tctgatecct	ttcctcttta	cggcacaaca	ttcatgtttg	acagaacatg	300
ctggaatgca	attgtttgca	acacogaagg	atttcctgcy	gtcgcctctt	cagtaggaag	360
cactgcattg	gtgataggac	acggtaattt	gattcacatt	taacttgcta	gttagtgata	420
aggggtggta	cacctgtttg	gtaaaatgag	aagcctcgga	aacttgggag	cttctctcct	480
accactaatg	gggagggcag	attattactg	ggattttctc	tgggggtgaat	taatttcaag	540
ccctaattgc	tgaaattccc	ctnggcaggc	tccagttttc	tcaactgcat	tgcaaaattc	600
cccc						604

<210> 91

<211> 858

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(858)

<223> n = A,T,C or G

<400> 91

tttttttttt	ttttttttta	tgattattat	tttttttatt	gatctttaca	tcctcagtgt	60
tggcagagtt	tctgatgctt	aataaacatt	tgttctgac	agataagtgg	aaaaaattgt	120
catttccctta	ttcaagccat	gcttttctgt	gatattctga	tcctagttga	acatacagaa	180

ataaatgtct	aaaacagcac	ctcgattctc	gtctataaca	ggactaagtt	cactgtgatc	240
ttaaataagc	ttggctaaaa	tgggacatga	gtggaggtag	tcacacttca	gcgaagaaag	300
agaatctcct	gtataatctc	accaggagat	tcaacgaatt	ccaccacact	ggactagtgg	360
atcccccggg	ctgcaggaat	tcgatatcaa	gcttatcgat	accgtcgacc	tcgagggggg	420
gccccgtacc	caattcgccc	tatagtgagt	cgtattacgc	gcgctcactg	gcgctcgttt	480
tacaaacgtcg	tgactgggaa	aaccctggcg	ttaccctaact	taatcgctt	gcagcacatc	540
ccccttttcgc	cagctggcgt	aatagcgaa	agcccgccacc	gatcgccctt	ncaacagttg	600
cgcagcctga	atggcgaaatg	ggacgcgccc	tgtagcggcg	cattaaagcg	cggcnggggtg	660
tggnggntcc	cccacgtgac	cgntacactt	ggcagcgcct	tacgcgggtc	nttcgctttc	720
ttcccttctt	ttctcgacac	gttcgcgggg	tttccccggn	agctnttaat	cgggggnctc	780
cctttanggg	tncnaattaa	nggnttacng	gaccttngan	cccaaaaact	ttgattaggg	840
ggaaggtccc	cgaagggg					858

<210> 92

<211> 585

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)... (585)

<223> n = A,T,C or G

<400> 92

gttgaatctc	ctgggtgagat	tatacaggag	attctctttc	ttcgtgaag	tgtgactacc	60
tccactcatg	tcccatttta	gccaagctta	tttaagatca	cagtgaactt	agtcctgtta	120
tagacgagaa	tcgaggtgct	gttttagaca	tttatttctg	tatgttcaac	taggatcaga	180
atatcacaga	aaagcatggc	ttgaataagg	aaatgacaat	tttttccact	tatctgatca	240
gaacaaatgt	ttattaagca	tcagaaactc	tgccaacact	gaggatgtaa	agatcaataa	300
aaaaaataat	aatcatnann	naaanannan	nngaagggcg	gccgccaccg	cggtggagct	360
ccagcttttg	ttcccttttag	tgaggggttaa	ttgcgcgctt	ggcggttaatc	atgggtcatag	420
ctgtttcctg	tgtgaaattg	ttatccggct	cacaattccn	cncaacatac	gagccgggaa	480
gcntnangtg	taaaagcctg	gggggtgccta	attgagttag	ctnactcaca	ttaattgngt	540
tgcgtccac	ttgccgcgtt	ttccantccg	ggaaacctgt	tcgnc		585

<210> 93

<211> 567

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)... (567)

<223> n = A,T,C or G

<400> 93

cggcagtggt	gctgtctgcg	tgtccacctt	ggaatctggc	tgaactggct	gggaggacca	60
agactgcggc	tgggggtggc	anggaaggga	accgggggct	gctgtgaagg	atcttggaac	120
ttccctgtac	ccaccttccc	cttgcttcat	gtttgtanag	gaaccttgtg	ccggccaagc	180
ccagtttctt	tgtgtgatac	actaatgtat	ttgctttttt	tgggaaatan	anaaaaaatca	240
attaaattgc	tantgtttct	ttgaannnnn	nnnnnnnnnn	nnnnnnnggg	ggggncgccc	300
ccnccgngga	aacnccccct	tttgttccct	ttaattgaaa	ggttaattng	cncnctggc	360
gttaancnt	gggccaaanc	tngttneccg	tgntgaaatt	gttnatcccc	tcccaaattc	420
ccccccnncc	ttccaaaccc	ggaaancctn	annntgttna	anccgggggg	gttgccctaan	480
ngnaattnaa	ccnaaccccc	ntttaaatng	nntttgcn	ccacnngccc	cnctttccca	540

nttcggggaa aacctntcc gtgccca

567

<210> 94
 <211> 620
 <212> DNA
 <213> Homo sapien

 <220>
 <221> misc_feature
 <222> (1)... (620)
 <223> n = A,T,C or G

<400> 94
 actagtcaaa aatgctaaaa taatttggga gaaaatattt ttttaagtagt gttatagttt 60
 catgtttatc ttttattatg ttttgtgaag ttgtgtcttt tcactaatta cctatactat 120
 gccaatattt ctttatatct atccataaca tttatactac atttgaana naatatgcac 180
 gtgaaactta acactttata aggtaaaaat gaggtttcca anatttaata atctgatcaa 240
 gttcttggtta tttccaaata gaatggactt ggtctgttaa gggctaagga gaagaggaag 300
 ataagggttaa aagtgtgttaa tgaccaaaca ttctaaaaga aatgcaaaaa aaaagtttat 360
 tttcaagcct tcgaactatt taaggaaagc aaaatcattt cctaaatgca tatcatttgt 420
 gagaatttct cattaatatc ctgaatcatt catttcacta aggctcatgt tnactccgat 480
 atgtctctaa gaaagtacta tttcatggtc caaacctggt tgccatantt gggtaaaggc 540
 tttcccttaa gtgtgaaant atttaaaatg aaattttcct ctttttaaaa attctttana 600
 aggggttaagg gtgttgggga 620

<210> 95
 <211> 470
 <212> DNA
 <213> Homo sapien

 <220>
 <221> misc_feature
 <222> (1)... (470)
 <223> n = A,T,C or G

<400> 95
 ctgcaccttc tctgcacagc ggatgaacct tgagcagctg aagaccagaa aagccactat 60
 nactttntgc ttaattcang agcttacang attcttcaaa gagtgngtcc agcatccttt 120
 gaaacatgag ttcttaccag cagaagcaga cctttacccc accacctcag cttcaacagc 180
 agcaggtgaa acaaccctac cagcctccac ctnaggaaat atttgttccc acaaccaagg 240
 agccatgcca ctcaaagggt ccacaacctg naaacacaaa nattccagag ccagggtgta 300
 ccaagggtccc tgagccagggt ctgtaccaan gtccctgagc cagggtgtac caangtcctt 360
 gagccaggat gtaccaagggt ccctgancca ggttgtccaa ggtccctgag ccagggtaca 420
 ccaagggcct gngccaggca gcatcaangt ccttgaccaa ggcttatcaa 470

<210> 96
 <211> 660
 <212> DNA
 <213> Homo sapien

 <220>
 <221> misc_feature
 <222> (1)... (660)
 <223> n = A,T,C or G

<400> 96

tttttttttt	tttttttttt	ggaattaaaa	gcaatttaaat	gagggcagag	caggaaacat	60
gcattttcttt	tcatttgaat	cttcagatga	accctgagca	gccgaagacc	agaaaagcca	120
tgaagacttt	ctgcttaatt	caggggctta	caggattctt	cagagtgtgt	gtgaacaaaa	180
gctttatagt	acgtattttt	aggatacaaa	taagagagag	actatggctt	ggggtgagaa	240
tgtactgatt	acaaggtcta	cagacaatta	agacacagaa	acagatggga	agaggggtgnc	300
cagcatctgg	nggttggtt	ctcaagggtt	tgtctgtgca	ccaaattact	tctgcttggn	360
cttctgctga	gctgggcctg	gagtgaacct	tgaaggacat	ggctctggtg	cctttgtgta	420
gcctgncaca	ggaactttgg	tgtatccttg	ctcagggaact	ttgatggcac	ctggctcagg	480
aaacttgatg	aagccttggt	caagggacct	tgatgcttgc	tggctcaggg	accttggnngn	540
ancctgggct	canggacctt	tgncicaacc	ttggcttcaa	gggaccttg	gnacatcctg	600
gcnnagggac	ccttggncc	aacctggggc	tnagggacc	ctttggnntnc	nanccttggc	660

<210> 97

<211> 441

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(441)

<223> n = A,T,C or G

<400> 97

gggaccatac	anagtattcc	tctcttcaca	ccaggaccag	ccactgttgc	agcatgagtt	60
cccagcagca	gaagcagccc	tgcattccac	cccctcagct	tcagcagcag	cagggtgaac	120
agccttgcca	gcctccacct	caggaaccat	gcattcccaa	aaccaaggag	ccctgccacc	180
ccaaggtgcc	tgagccctgc	caccccaaag	tgcttgagcc	ctgccagccc	aaggttccag	240
agccatgcca	ccccaaggtg	cctgagccct	gcccttcaat	agtcactcca	gcaccagccc	300
agcagaanac	caagcagaag	taatgtggtc	cacagccatg	cccttgagga	gccggccacc	360
agatgctgaa	tccctatcc	cattctgtgt	atgagtccca	tttgccctgc	aattagcatt	420
ctgtctcccc	caaaaaaaaa	a				441

<210> 98

<211> 600

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(600)

<223> n = A,T,C or G

<400> 98

gtattcctct	cttcacacca	ggaccagcca	ctgttgagc	atgagttccc	agcagcagaa	60
gcagccctgc	atcccacccc	ctcagcttca	gcagcagcag	gtgaaacagc	cttgccagcc	120
tccacctcag	gaacctgca	tccccaaaac	caaggagccc	tgccaccca	aggtgacctga	180
gcctgtccac	cccaaagtgc	ctgagccctg	ccagcccaaag	gttccagagc	catgccaccc	240
caaggtgcct	gagccctgcc	cttcaatagt	cactccagca	ccagcccagc	agaanaccaa	300
gcagaagtaa	tgtgggtccac	agccatgccc	ttgaggagcc	ggccaccana	tgctgaatcc	360
cctatcccat	tctgtgtatg	agtcacattt	gccttgcaat	tagcattctg	tctcccccaa	420
aaaagaatgt	gctatgaagc	tttctttcct	acacactctg	agtctctgaa	tgaagctgaa	480
ggtcttaant	acaganctag	ttttcagctg	ctcagaattc	tctgaagaaa	agatttaaga	540
tgaaaggcaa	atgattcagc	tccttattac	ccattaaat	tcnctttcaa	ttccaaaaaa	600

<210> 99
 <211> 667
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (667)
 <223> n = A,T,C or G

<400> 99
 actagtgact gagttcctgg caaagaaatt tgacctggac cagttgataa ctcatgtttt 60
 accattttaa aaaatcagtg aaggatttga gctgctcaat tcaggacaaa gcattcgaac 120
 ggtcctgacg ttttgagatc caaagtggca ggaggtctgt gttgtcatgg tgaactggag 180
 tttctcttgt gagagttccc tcatctgaaa tcatgtatct gtctcacaaa tacaagcata 240
 agtgaagat ttgttgaaga catagaaccc ttataaagaa ttattaacct ttataaacat 300
 ttaaagtctt gtgagcacct gggaattagt ataataacaa tgtnnatatt tttgatttac 360
 attttgtaag gctataattg tatcttttaa gaaaacatac cttggatttc tatgttgaaa 420
 tggagatttt taagagtttt aaccagctgc tgcagatata ttactcaaaa cagatatagc 480
 gtataaagat atagtaaag catctcctag agtaatatc acttaacaca ttggaaacta 540
 ttatttttta gatttgaata tnaatgttat tttttaaaca cttgttatga gttacttggg 600
 attacatttt gaaatcagtt cattccatga tgcanattac tgggattaga ttaagaaaga 660
 cggaaaa 667

<210> 100
 <211> 583
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (583)
 <223> n = A,T,C or G

<400> 100
 gttttgtttg taagatgac acagtcattg tacactgatc taaaggacat atatataacc 60
 ctttaaaaaa aaaatcactg cctcattctt atttcaagat gaatttctat acagactaga 120
 tgtttttctg aagatcaatt agacattttg aaaatgattt aaagtgtttt ctttaatgtt 180
 ctctgaaaac aagtttcttt ttagtattta accaaaaaag tgcccttttt gtcactggat 240
 tctcctagca ttcattgattt ttttttcata caatgaaatt aaaattgcta aaatcatgga 300
 ctggctttct ggttggattt caggtaagat gtgtttaagg ccagagcttt tctcagtatt 360
 tgattttttt cccaatatt tgatttttta aaaatatata catnggtgct gcatttatat 420
 ctgctggttt aaaattctgt catatttcac ttctagcctt ttagttatgg caaatcatat 480
 ttacttttta cttaaagcat ttggttnattt ggantatctg gttctannct aaaaaanta 540
 attctatnaa ttgaantttt ggtactcnnc catatttggg tcc 583

<210> 101
 <211> 592
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (592)
 <223> n = A,T,C or G

<400> 101

gtggagacgt	acaaagagca	gccgctcaag	acacctggga	agaaaaagaa	aggcaagccc	60
gggaaacgca	aggagcagga	aaagaaaaaa	cggcgaactc	gctctgcctg	gtagactct	120
ggagtgactg	ggagtgggct	agaaggggac	cacctgtctg	acacctccac	aacgtcgctg	180
gagctcgatt	cacggaggca	ttgaaatttt	cagcaganac	cttccaagga	catattgcag	240
gattctgtaa	tagtgaacat	atggaaaagta	ttagaaatat	ttattgtctg	taaatactgt	300
aaatgcattg	gaataaaaact	gtctcccca	ttgctctatg	aaactgcaca	ttggtcattg	360
tgaatatttt	tttttttgcc	aaggctaata	caattattat	tatcacattt	accataattt	420
attttgtcca	ttgatgtatt	tattttgtaa	atgtatcttg	gtgctgctga	atttctatat	480
tttttgtaca	taatgcnttt	anatatacct	atcaagtttg	ttgataaatg	acncaatgaa	540
gtgncncnan	ttgngggtg	aatttaatga	atgcctaatt	ttattatccc	aa	592

<210> 102

<211> 587

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (587)

<223> n = A,T,C or G

<400> 102

cgctctaagc	acttagacta	catcagggaa	gaacacagac	cacatccctg	tcctcatgcg	60
gcttatgttt	tctggaagaa	agtggagacc	nagtccttgg	ctttagggct	ccccggctgg	120
gggctgtgca	ntccggtcag	ggcgggaagg	gaaatgcacc	gctgcatgtg	aacttacagc	180
ccaggcggat	gcccttccc	ttagcactac	ctggcctcct	gcacccctc	gcctcatgtt	240
cctcccacct	tcaanaaatg	aanaaccca	tgggccagc	cccttgccct	ggggaaccaa	300
ggcagccttc	caaaactcag	gggctgaagc	anactattag	ggcaggggct	gactttgggt	360
gacactgccc	attccctctc	agggcagctc	angtcacccn	ggncctctga	accagcctg	420
ttcctttgaa	aaagggcaaa	actgaaaagg	gcttttcta	naaaaagaaa	aaccagggaa	480
ctttgccagg	gcttcnntnt	taccaaaacn	ncttctcnng	gatttttaat	tccccattng	540
gcctccactt	accnggggcn	atgccccaaa	attaanaatt	tcccatc		587

<210> 103

<211> 496

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (496)

<223> n = A,T,C or G

<400> 103

anaggactgg	ccctacntgc	tctctctcgt	cctacctatc	aatgcccaac	atggcagaac	60
ctgcancctt	tgncaactgc	anatggaaac	ctctcagtgt	cttgacatca	ccctaccnt	120
gcggtgggtc	tccaccacaa	ccactttgac	tctgtggtcc	ctgnanggtg	gnttctcctg	180
actggcagga	tggaccttan	ccnacatata	cctctgttcc	ctctgctnag	anaaagaatt	240
cccttaacat	gatataatcc	acccatgcaa	ntngetactg	gccagctac	catttaccat	300
ttgcctacag	aatttcattc	agtctacact	ttggcattct	ctctggcgat	agagtgtggc	360
tgggctgacc	gcaaaagggtg	ccttacacac	tggcccccac	cctcaaccgt	tgacncatca	420
gangcttgcc	tcctccttct	gattnncccc	catgttggat	atcagggtgc	tcnagggtt	480
ggaaaagaaa	caaaac					496

<210> 104
 <211> 575
 <212> DNA
 <213> Homo sapien

 <220>
 <221> misc_feature
 <222> (1) ... (575)
 <223> n = A,T,C or G

<400> 104
 gcacctgctc tcaatccnnc tctcaccatg atcctccgcc tgcanaaact cctctgccaa 60
 ctatggangt ggtttcnggg gtggctcttg ccaactggga agaagccgtg gtgtctctac 120
 ctgttcaact cngtttgtgt ctgggggatc aactnngggc tatggaagcg gctnaactgt 180
 tgttttggtg gaagggctgg taattggctt tgggaagtng cttatngaag ttggcctngg 240
 gaagttgcta ttgaaagtng ccntggaagt ngntttggtg gggggttttg ctggtggcct 300
 ttgttnaatt tgggtgcttt gtnaatggcg gccccctcnc ctgggcaatg aaaaaaatca 360
 ccnatgcngn aaacctcnac nnaacagcct gggcttcctt cacctcgaaa aaagttgctc 420
 ccccccaaaa aaaggncaan cccctcaann tggaangttg aaaaaatcct cgaatgggga 480
 nccnnaaaac aaaaancccc ccntttcccn gnaanggggg aaataccncc cccccactta 540
 cnaaaaccct tntaaaaaac cccccgggaa aaaaa 575

<210> 105
 <211> 619
 <212> DNA
 <213> Homo sapien

 <220>
 <221> misc_feature
 <222> (1) ... (619)
 <223> n = A,T,C or G

<400> 105
 cactagtagg atagaaacac tgtgtcccgga gagtaaggag agaagctact attgattaga 60
 gcctaaccce ggtaactgc aagaagaggc gggatacttt cagctttcca tgtaactgta 120
 tgcataaagc caatgtagtc cagtttctaa gatcatgttc caagctaact gaatccact 180
 tcaatacaca ctcatgaact cctgatggaa caataacagg cccaagcctg tggatgatg 240
 tgcacacttg ctgactcan aaaaaatact actctcataa atgggtggga gtattttggt 300
 gacaacctac tttgcttggc tgagtgaagg aatgatattc atatattcat ttattccatg 360
 gacatttagt tagtgctttt tatataccag gcatgatget gagtgacact cttgtgtata 420
 tttccaaatt tttgtacagt cgctgcacat atttgaaatc atatattaag acttccaaaa 480
 aatgaagtcc ctggtttttc atggcaactt gatcagtaaa ggattcnctt ctggttggtg 540
 cttaaaacat ctactatatn gttanatatga aattcctttt ccccnctcc cgaaaaaana 600
 aagtgggtgg gaaaaaaa 619

<210> 106
 <211> 506
 <212> DNA
 <213> Homo sapien

 <220>
 <221> misc_feature
 <222> (1) ... (506)
 <223> n = A,T,C or G

<400> 106

cattggtnct	ttcatttgct	ntggaagtgt	nmatctctaa	cagtggacaa	agttcccngt	60
gccttaaact	ctgtnacact	tttggaant	gaaaantng	tantatgata	ggttattctg	120
angtanagat	gttctggata	ccattanatn	tgccccngt	gtcagagget	catattgtgt	180
tatgtaaag	gtatntcatt	cgctactatn	antcaatng	aaatanggtc	tttgggttat	240
gaatantng	cagcncanct	nanangctgt	ctgtngtatt	cattgtggc	atagcacctc	300
acancattgt	aacctcnatc	nagtggagaca	nactagnaana	ttcctagtga	tggctcanga	360
ttccaaatgg	ntcatntcn	aatgtttaaa	agttanttaa	gtgtaagaaa	tacagactgg	420
atgttccacc	aactagtacc	tgtaatgacn	ggcctgtccc	aacacatctc	ccttttccat	480
gactgtggta	ncccgcatcg	gaaaaa				506

<210> 107

<211> 452

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)... (452)

<223> n = A,T,C or G

<400> 107

gttgagtctg	tactaaacag	taagatatct	caatgaacca	taaattcaac	tttgtaaaaa	60
tcttttgaag	catagataat	attgtttggg	aaatgtttct	tttgtttggg	aaatgtttct	120
tttaaagacc	ctcctattct	ataaaactct	gcatgtagag	gcttgtttac	ccttctctct	180
ctaaggttta	caataggagt	ggtgatttga	aaaatataaa	attatgagat	tgggttttct	240
gtggcataaa	ttgcatcact	gtatcatttt	cctttttaac	cggttaagant	ttcagtttgt	300
tggaaagtaa	ctgtganaac	ccagtttccc	gtccatctcc	cttagggact	acccatagaa	360
catgaaaagg	tccccacnga	agcaagaaga	taagtctttc	atggctgctg	gttgcttaaa	420
ccacttttaa	accaaaaaat	tccccttgga	aa			452

<210> 108

<211> 502

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)... (502)

<223> n = A,T,C or G

<400> 108

atcttcttcc	cttaattagt	tnttatttat	ntattaaatt	ttattgcatg	tcttggaaca	60
caaaaagaga	ttgtagattg	gcttctggct	ccccaaaagc	ccataacaga	aagtaccaca	120
agaccncaac	tgaagcttaa	aaaatctatc	acatgtataa	tacctttnga	agaacattaa	180
tanagcatat	aaaactttta	acatntgctt	aatgttgtn	aattataaaa	ntaatngaaa	240
aaaatgtccc	tttaacatnc	aatatcccac	atagtgttat	ttnaggggat	taccnngnaa	300
naaaaaaagg	gtagaaggga	tttaatgaaa	actctgcttn	ccatttctgt	ttanaaacgt	360
ctccagaaca	aaaacttntc	aantctttca	gctaaccgca	tttgagctna	ggccactcaa	420
aaactccatt	agncccactt	tctaanggtc	tctanagctt	actaancctt	ttgaccctt	480
accctggnta	ctcctgcctt	ca				502

<210> 109

<211> 1308

<212> DNA

<213> Homo sapien

<400> 109

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ggcatcttga ctgcaattgg catggtcctc ctggggaccc gaggagccac cgcttcccag      180
ttggaggagg tgtttctactc tgaaaaagag acgaagagct caagaataaa ggctgaagaa      240
aaagaggtga ttgagaacac agaagcagta catcaacaat tccaaaagtt ttgactgaa      300
ataagcaaac tctaataatga ttatgaactg aacataacca acaggctgtt tggagaaaaa      360
acatacctct tccttcaaaa atacttagat tatgttgaaa aatattatca tgcattctctg      420
gaacctgttg attttgtaaa tgcagccgat gaaagtcgaa agaagattaa ttcttgggtt      480
gaaagcaaaa caaatgaaaa aatcaaggac ttgttcccag atggctctat tagtagctct      540
accaagctgg tgctggtgaa catggtttat tttaaagggc aatgggacag ggagttaaag      600
aaagaaaata ctaaggaaga gaaattttgg atgaataaga gcacaagtaa atctgtacag      660
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gatggcctgg agaagataat agataaaata agtcttgaga aattggtaga gtggactagt      840
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agttacgata tagaggcggc cctggctgcc atggggatgg gcgatgcctt cagtgcacac      960
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tttactgtca catccgcccc aggtcatgaa aatgttcaact gcaatcatcc cttcctgttc      1140
ttcatcaggc acaatgaatc caacgacatc ctcttcttcg gcagattttc ttctccttaa      1200
gatgategtt gccatggcat tgctgctttt agcaaaaaac aactaccagt gttactcata      1260
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<210> 110

<211> 391

<212> PRT

<213> Homo sapien

<400> 110

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          20           25           30
Gly Ile Leu Thr Ala Ile Gly Met Val Leu Leu Gly Thr Arg Gly Ala
          35           40           45
Thr Ala Ser Gln Leu Glu Glu Val Phe His Ser Glu Lys Glu Thr Lys
          50           55           60
Ser Ser Arg Ile Lys Ala Glu Glu Lys Glu Val Ile Glu Asn Thr Glu
          65           70           75           80
Ala Val His Gln Gln Phe Gln Lys Phe Leu Thr Glu Ile Ser Lys Leu
          85           90           95
Thr Asn Asp Tyr Glu Leu Asn Ile Thr Asn Arg Leu Phe Gly Glu Lys
          100          105          110
Thr Tyr Leu Phe Leu Gln Lys Tyr Leu Asp Tyr Val Glu Lys Tyr Tyr
          115          120          125
His Ala Ser Leu Glu Pro Val Asp Phe Val Asn Ala Ala Asp Glu Ser
          130          135          140
Arg Lys Lys Ile Asn Ser Trp Val Glu Ser Lys Thr Asn Glu Lys Ile
          145          150          155          160
Lys Asp Leu Phe Pro Asp Gly Ser Ile Ser Ser Ser Thr Lys Leu Val
          165          170          175

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Leu Val Asn Met Val Tyr Phe Lys Gly Gln Trp Asp Arg Glu Phe Lys
 180 185 190
 Lys Glu Asn Thr Lys Glu Glu Lys Phe Trp Met Asn Lys Ser Thr Ser
 195 200 205
 Lys Ser Val Gln Met Met Thr Gln Ser His Ser Phe Ser Phe Thr Phe
 210 215 220
 Leu Glu Asp Leu Gln Ala Lys Ile Leu Gly Ile Pro Tyr Lys Asn Asn
 225 230 235 240
 Asp Leu Ser Met Phe Val Leu Leu Pro Asn Asp Ile Asp Gly Leu Glu
 245 250 255
 Lys Ile Ile Asp Lys Ile Ser Pro Glu Lys Leu Val Glu Trp Thr Ser
 260 265 270
 Pro Gly His Met Glu Glu Arg Lys Val Asn Leu His Leu Pro Arg Phe
 275 280 285
 Glu Val Glu Asp Ser Tyr Asp Leu Glu Ala Val Leu Ala Ala Met Gly
 290 295 300
 Met Gly Asp Ala Phe Ser Glu His Lys Ala Asp Tyr Ser Gly Met Ser
 305 310 315 320
 Ser Gly Ser Gly Leu Tyr Ala Gln Lys Phe Leu His Ser Ser Phe Val
 325 330 335
 Ala Val Thr Glu Glu Gly Thr Glu Ala Ala Ala Thr Gly Ile Gly
 340 345 350
 Phe Thr Val Thr Ser Ala Pro Gly His Glu Asn Val His Cys Asn His
 355 360 365
 Pro Phe Leu Phe Phe Ile Arg His Asn Glu Ser Asn Ser Ile Leu Phe
 370 375 380
 Phe Gly Arg Phe Ser Ser Pro
 385 390

<210> 111

<211> 1419

<212> DNA

<213> Homo sapien

<400> 111

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 ggcgcgctca gcactcgact tgggtttgat cttttcaaag agctgaagaa aacaaatgat 180
 ggcaacatct tcttttcccc tgtgggcac ttgactgcaa ttggcatggg cctcctgggg 240
 acccgaggag ccaccgcttc ccagttggag gaggtgttct actctgaaaa agagacgaag 300
 agctcaagaa taaaggctga agaaaaagag gtggttaagaa taaaggctga aggaaaagag 360
 attgagaaca cagaagcagt acatcaacaa ttccaaaagt ttttgactga aataagcaaa 420
 ctactaatg attatgaact gaacataacc aacaggctgt ttggagaaaa aacatacctc 480
 ttccttcaaa aatacttaga ttatgttgaa aaatattatc atgcatctct ggaacctgtt 540
 gattttgtaa atgcagcoga tgaaagtcga aagaagatta attcctgggt tgaaagcaaa 600
 acaaatgaaa aaatcaagga cttgttccca gatggctcta ttagtagctc taccaagctg 660
 gtgctggtga acatggttta ttttaaaggg caatgggaca gggagttaa gaaagaaaat 720
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 ccatataaaa acaacgacct aagcatgttt gtgcttctgc ccaacgacat cgatggcctg 900
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acatccgccc caggatcatga aaatgttcac tgcaatcatc ccttctctgtt cttcatcagg 1260
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<210> 112

<211> 400

<212> PRT

<213> Homo sapien

<400> 112

Met	Asp	Ser	Leu	Gly	Ala	Val	Ser	Thr	Arg	Leu	Gly	Phe	Asp	Leu	Phe	1	5	10	15
Lys	Glu	Leu	Lys	Lys	Thr	Asn	Asp	Gly	Asn	Ile	Phe	Phe	Ser	Pro	Val	20	25	30	
Gly	Ile	Leu	Thr	Ala	Ile	Gly	Met	Val	Leu	Leu	Gly	Thr	Arg	Gly	Ala	35	40	45	
Thr	Ala	Ser	Gln	Leu	Glu	Glu	Val	Phe	His	Ser	Glu	Lys	Glu	Thr	Lys	50	55	60	
Ser	Ser	Arg	Ile	Lys	Ala	Glu	Glu	Lys	Glu	Val	Val	Arg	Ile	Lys	Ala	65	70	75	80
Glu	Gly	Lys	Glu	Ile	Glu	Asn	Thr	Glu	Ala	Val	His	Gln	Gln	Phe	Gln	85	90	95	
Lys	Phe	Leu	Thr	Glu	Ile	Ser	Lys	Leu	Thr	Asn	Asp	Tyr	Glu	Leu	Asn	100	105	110	
Ile	Thr	Asn	Arg	Leu	Phe	Gly	Glu	Lys	Thr	Tyr	Leu	Phe	Leu	Gln	Lys	115	120	125	
Tyr	Leu	Asp	Tyr	Val	Glu	Lys	Tyr	Tyr	His	Ala	Ser	Leu	Glu	Pro	Val	130	135	140	
Asp	Phe	Val	Asn	Ala	Ala	Asp	Glu	Ser	Arg	Lys	Lys	Ile	Asn	Ser	Trp	145	150	155	160
Val	Glu	Ser	Lys	Thr	Asn	Glu	Lys	Ile	Lys	Asp	Leu	Phe	Pro	Asp	Gly	165	170	175	
Ser	Ile	Ser	Ser	Ser	Thr	Lys	Leu	Val	Leu	Val	Asn	Met	Val	Tyr	Phe	180	185	190	
Lys	Gly	Gln	Trp	Asp	Arg	Glu	Phe	Lys	Lys	Glu	Asn	Thr	Lys	Glu	Glu	195	200	205	
Lys	Phe	Trp	Met	Asn	Lys	Ser	Thr	Ser	Lys	Ser	Val	Gln	Met	Met	Thr	210	215	220	
Gln	Ser	His	Ser	Phe	Ser	Phe	Thr	Phe	Leu	Glu	Asp	Leu	Gln	Ala	Lys	225	230	235	240
Ile	Leu	Gly	Ile	Pro	Tyr	Lys	Asn	Asn	Asp	Leu	Ser	Met	Phe	Val	Leu	245	250	255	
Leu	Pro	Asn	Asp	Ile	Asp	Gly	Leu	Glu	Lys	Ile	Ile	Asp	Lys	Ile	Ser	260	265	270	
Pro	Glu	Lys	Leu	Val	Glu	Trp	Thr	Ser	Pro	Gly	His	Met	Glu	Glu	Arg	275	280	285	
Lys	Val	Asn	Leu	His	Leu	Pro	Arg	Phe	Glu	Val	Glu	Asp	Ser	Tyr	Asp	290	295	300	
Leu	Glu	Ala	Val	Leu	Ala	Ala	Met	Gly	Met	Gly	Asp	Ala	Phe	Ser	Glu	305	310	315	320
His	Lys	Ala	Asp	Tyr	Ser	Gly	Met	Ser	Ser	Gly	Ser	Gly	Leu	Tyr	Ala	325	330	335	
Gln	Lys	Phe	Leu	His	Ser	Ser	Phe	Val	Ala	Val	Thr	Glu	Glu	Gly	Thr	340	345	350	

Glu Ala Ala Ala Ala Thr Gly Ile Gly Phe Thr Val Thr Ser Ala Pro
 355 360 365
 Gly His Glu Asn Val His Cys Asn His Pro Phe Leu Phe Phe Ile Arg
 370 375 380
 His Asn Glu Ser Asn Ser Ile Leu Phe Phe Gly Arg Phe Ser Ser Pro
 385 390 395 400

<210> 113
 <211> 957
 <212> DNA
 <213> Homo sapien

<400> 113

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gaaacatgag	ttcttaccag	cagaagcaga	cctttacccc	accacctcag	cttcaacagc	180
agcaggtgaa	acaacccagc	cagcctccac	ctcaggaaat	atttgttccc	acaaccaagg	240
agccatgcc	ctcaaagggt	ccacaacctg	gaaacacaaa	gattccagag	ccaggctgta	300
ccaaggtccc	tgagccaggc	tgtaccaagg	tccttgagcc	aggttgtagc	aaggtccctg	360
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ccaaggtccc	tgagccaggc	agcatcaagg	tccttgacca	aggttctatc	aagtttcctg	480
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ccctcttccc	atctgtttct	gtgtcttaat	tgtctgtaga	ccttgtaatc	agtacattct	720
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<210> 114
 <211> 161
 <212> PRT
 <213> Homo sapien

<400> 114

Met	Ser	Ser	Tyr	Gln	Gln	Lys	Gln	Thr	Phe	Thr	Pro	Pro	Pro	Gln	Leu
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Gln	Gln	Gln	Gln	Val	Lys	Gln	Pro	Ser	Gln	Pro	Pro	Pro	Pro	Gln	Ile
				20					25					30	
Phe	Val	Pro	Thr	Thr	Lys	Glu	Pro	Cys	His	Ser	Lys	Val	Pro	Gln	Pro
				35					40					45	
Gly	Asn	Thr	Lys	Ile	Pro	Glu	Pro	Gly	Cys	Thr	Lys	Val	Pro	Glu	Pro
				50					55					60	
Gly	Cys	Thr	Lys	Val	Pro	Glu	Pro	Gly	Cys	Thr	Lys	Val	Pro	Glu	Pro
				65					70					75	
Gly	Cys	Thr	Lys	Val	Pro	Glu	Pro	Gly	Cys	Thr	Lys	Val	Pro	Glu	Pro
				85					90					95	
Gly	Tyr	Thr	Lys	Val	Pro	Glu	Pro	Gly	Ser	Ile	Lys	Val	Pro	Asp	Gln
				100					105					110	
Gly	Phe	Ile	Lys	Phe	Pro	Glu	Pro	Gly	Ala	Ile	Lys	Val	Pro	Glu	Gln
				115					120					125	
Gly	Tyr	Thr	Lys	Val	Pro	Val	Pro	Gly	Tyr	Thr	Lys	Val	Pro	Glu	Pro
				130					135					140	
Cys	Pro	Ser	Thr	Val	Thr	Pro	Gly	Pro	Ala	Gln	Gln	Lys	Thr	Lys	Gln

145
Lys

150

155

160

<210> 115
 <211> 506
 <212> DNA
 <213> Homo sapien

 <220>
 <221> misc_feature
 <222> (1)... (506)
 <223> n = A,T,C or G

<400> 115

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gccttaaaact	ctgtnacact	tttgggaant	gaaaanttng	tantatgata	ggttattctg	120
angtanagat	gttctggata	ccattanatn	tgccccngt	gtcagaggct	catatttgtt	180
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gaatantnng	cagcncanct	nanangctgt	ctgtngtatt	cattgtggtc	atagcacctc	300
acancattgt	aacctcmtc	nagtggagaca	nactagnaana	ttcctagtga	tggtccanga	360
ttccaaatgg	ntcatntcn	aatgtttaaa	agttanttaa	gtgtaagaaa	tacagactgg	420
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gactgtggta	ncccgcatcg	gaaaaa				506

<210> 116
 <211> 3079
 <212> DNA
 <213> Homo sapien

<400> 116

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<210> 117

<211> 6921

<212> DNA

<213> Homo sapien

<400> 117

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8948

<210> 120
 <211> 587
 <212> DNA
 <213> Homo sapien

 <220>
 <221> misc_feature
 <222> (1) ... (587)
 <223> n = A,T,C or G

<400> 120
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 gggctgtgca ntccggtcag ggcgggaagg gaaatgcacc gctgcatgtg aacttacagc 180
 ccaggcggat gccccttccc ttagcaactac ctggcctcct gcacccctc gcctcatgtt 240
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 gacactgccc attcctctc agggcagctc angtcacccn ggmctcttga acccagcctg 420
 ttcttttgaa aaagggcaaa actgaaaagg gcttttctta naaaaagaaa aaccagggaa 480
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<210> 121
 <211> 619
 <212> DNA
 <213> Homo sapien

 <220>
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 <222> (1) ... (619)
 <223> n = A,T,C or G

<400> 121
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 tgcataaagc caatgtagtc cagtttctaa gatcatgttc caagctaact gaatccact 180
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 aatgaagtcc ctggtttttc atggcaactt gatcagtaaa ggattcnct ctggttggt 540
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 aagtgggtgg gaaaaaaa 619

<210> 122
 <211> 1475
 <212> DNA
 <213> Homo sapien

<400> 122
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<210> 123

<211> 2294

<212> DNA

<213> Homo sapien

<400> 123

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<210> 124

<211> 956

<212> DNA

<213> Homo sapien

<400> 124

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<210> 125

<211> 486

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(486)

<223> n = A,T,C or G

<400> 125

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agcgcaggtt	ttggatacta	gagaaagtca	tttgcttgta	ctattgccat	tttagaaagc	420
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<210> 126

<211> 3552

<212> DNA

<213> Homo sapien

<400> 126

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<210> 127

<211> 754

<212> DNA

<213> Homo sapien

<400> 127

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<210> 128

<211> 374

<212> DNA

<213> Homo sapien

<400> 128

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<210> 129

<211> 546

<212> DNA

<213> Homo sapien

<400> 129

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<210> 130

<211> 5156

<212> DNA

<213> Homo sapien

<400> 130

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<210> 131

<211> 671

<212> DNA

<213> Homo sapien

<400> 131

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<210> 132

<211> 590

<212> DNA

<213> Homo sapien

<400> 132

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<210> 133

<211> 581

<212> DNA

<213> Homo sapien

<400> 133

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<210> 134

<211> 4797

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (4797)

<223> n = A,T,C or G

<400> 134

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tgacaacggt	tcagcgactc	cgttggccac	tcogagagtg	ggccagtctg	tggatcagag	4560
atgcaccacc	aagccaaggg	aacctgtgtc	cgttattcga	tactgcgact	ttctgcctgg	4620
agtgtatgac	tgcacatgac	tcgggggtgg	ggaaaggggt	cggctgacca	tgctcatctg	4680
ctggtccgtg	ggacggtncc	caagccagag	gtgggttcac	ttgtgtaacg	acaataaacg	4740
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<210> 135

<211> 2856

<212> DNA

<213> Homo sapien

<400> 135

tagtcgcggg	tccccgagtg	agcacgccag	ggagcaggag	accaaacgac	gggggtcgga	60
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cgcacgcccg	tcgccacccg	cgtaccgggc	gcagccagag	ccaccagcgc	agcgtgcca	180
tggagcccag	cagcaagaag	ctgacgggtc	gcctcatgct	ggctgtggga	ggagcagtgc	240
ttggctccct	gcagtttggc	tacaacactg	gagtcataca	tgccccccag	aagggtgatcg	300
aggagtctta	caaccagaca	tgggtccacc	gctatgggga	gagcatcctg	cccaccacgc	360
tcaccacgct	ctggtccctc	tcagtggcca	tcttttctgt	tgggggcatg	attggctcct	420
tctctgtggg	ccttttctgt	aacgcctttg	gcccggcgaa	ttcaatgctg	atgatgaacc	480
tgctggcctt	cgtgtccgcc	gtgctcatgg	gcttctcgaa	actgggcaag	tcctttgaga	540
tgctgatcct	gggcccgttc	atcatcggtg	tgtactgcgg	cctgaccaca	ggcttcgtgc	600
ccatgtatgt	gggtgaagtg	tcaccacacg	cctttcgtgg	ggccctgggc	accctgcacc	660
agctgggcat	cgtcgtcggc	atcctcatcg	cccagggtgt	cggcctggac	tccatcatgg	720
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tataaatggc	tgttttttag	aaacatggtt	ttgaaatgct	tgtggattga	gggtaggagg	2580
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tttgatccct	gttaccacaga	gaatatatac	attctttatc	ttgacattca	aggcatttct	2760
atcacatatt	tgatagttgg	tgttcaaaaa	aacactagtt	ttgtgccagc	cgtgatgctc	2820
aggcttgaaa	tcgcattatt	ttgaatgtga	agggaa			2856

<210> 136

<211> 356

<212> DNA

<213> Homo sapien

<400> 136

gggtggagcca	aatgaagaaa	atgaagatga	aagagacaga	cacctcagtt	tttctggatc	60
aggcattgat	gatgatgaag	attttatctc	cagcaccatt	tcaaccacac	cacgggcttt	120
tgaccacaca	aaacagaacc	aggactggac	tcagtggaac	ccaagccatt	caaaccggga	180
agtgtacttt	cagacaacca	caaggatgac	tgatgtagac	agaaatggca	ccactgctta	240
tgaaggaaac	tggaaaccag	aagcacaccc	tcccctcatt	caccatgagc	atcatgagga	300
agaagagacc	ccacattcta	caagcacaat	ccaggcaact	cctagtagta	caacgg	356

<210> 137

<211> 356

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (356)

<223> n = A,T,C or G

<400> 137

gcaggtggag	aagacatttt	attgttcctg	gggtctctgg	aggeccattg	gtggggctgg	60
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gtcactggct	gccccggaa	cagggcgctg	ctccatggct	ctgcttggtg	tagtctgtgg	120
ctatgtctcc	cagcaaggac	agaaactcag	aaaaatcaat	cttcttatcc	tcattcttgt	180
cctttttctc	aaagacatcg	gcgaggtaat	ttgtgccctt	tttacctcgg	cccgcgacca	240
cgctaaggcc	aaanttcag	acanayggcc	gggcccgtnc	nataggggan	cccaacttgg	300
ggacccaaac	tctggcgcg	aaacacangg	gcataagctt	gnttcctgtg	gggaaa	356

<210> 138

<211> 353

<212> DNA

<213> Homo sapien

<400> 138

aggtccagtc	ctccacttgg	cctgatgaga	gtggggagtg	gcaagggacg	tttctcctgc	60
aatagacact	tagatttctc	tcttggtgga	agaaaccacc	tgtccatcca	ctgactcttc	120
tacattgatg	tggaaattgc	tgctgctacc	accacctcct	gaagaggctt	ccctgatgcc	180
aatgccagcc	atcttggcat	cctggccctc	gagcaggctg	cggtaaagtag	cgatctcctg	240
ctccagcgt	gtctttatgt	caagcagcat	cttgtaactcc	tggttctgag	cctccatctc	300
gcatcggagc	tcactcagac	ctcgscgsg	mssmcgtam	gccgaattcc	agc	353

<210> 139

<211> 371

<212> DNA

<213> Homo sapien

<400> 139

agcgtggctg	cggccgaggt	ccatccgaag	caagattgca	gatggcagtg	tgaagagaga	60
agacatattc	tacacttcaa	agctttgggtg	caattcccat	cgaccagagt	tggtccgacc	120
agccttgga	aggtcactga	aaaatcttca	attggattat	gttgacctct	accttattca	180
ttttccagtg	tctgtaaagc	caggtgagga	agtgatccca	aaagatgaaa	atggaaaaat	240
actatttgac	acagtggatc	tctgtgccac	gtgggaggcc	gtggagaagt	gtaaagatgc	300
aggattggac	ctgcccgggc	ggccgctcga	aagccgaatt	ccagcacact	ggcggccggt	360
actagtggat	c					371

<210> 140

<211> 370

<212> DNA

<213> Homo sapien

<400> 140

tagcgtggtc	gcggccgagg	tccatctccc	tttgggaact	agggggctgc	tggtgggaaa	60
tgggagccag	ggcagatgtt	gcattccttt	gtgtccctgt	aaatgtggga	ctacaagaag	120
aggagctgcc	tgagtggtag	tttctcttcc	tggtaatcct	ctggcccagc	ctcatggcag	180
aatagaggta	tttttaggct	atttttgtaa	tatggcttct	gggtcaaaatc	cctgtgtagc	240
tgaattccca	agccctgcat	tgtacagccc	cccactcccc	tcaccaccta	ataaaggaat	300
agttaacact	caaaaaaaaa	aaaaaacctg	cccgggcggc	cgctcgaaag	ccgaattcca	360
gcacactggc						370

<210> 141

<211> 371

<212> DNA

<213> Homo sapien

<400> 141

tagcgtggtc	gcggccgagg	tcctctgtgc	tgccgtgcac	agcccgatgg	taccagcgca	60
gggtgtaggc	agtgcaggag	ccctcatcca	gtggcaggga	acaggggtca	tcactatccc	120

aaggagcttc agggtcctgg tactcctcca	cagaatactc ggagtattca	gagtactcat	180
catcctcagg gggtagccgc tcttctctct	ctgcatgaga gacgcggagc	acaggcacag	240
catggagctg ggagccggca gtgtctgcag	cataactagg gaggggtcgt	gatccagatg	300
cgatgaactg gccctggcag gcacagtgt	gactcatctc ttggcgacct	gcccgggcgg	360
ccgctogaag c			371

<210> 142

<211> 343

<212> DNA

<213> Homo sapien

<400> 142

gcgttttgag gccaatggtg taaaaggaaa	tatcttcaca taaaaactag	atggaagcat	60
tgtcagaaac ctctttgtga tgtttgcttt	caactcacag agttgaacat	tccttttcat	120
agagcagttt tgaacactc tttttagaa	tttgcaagcg gatgattgga	tcgctatgag	180
gtcttcattg gaaacgggat acctttacat	aaaaactaga cagtagcatt	ctcagaaatt	240
tctttgggat gtgggcattc aaccacaga	ggagaacttc atttgataga	gcagttttga	300
aacacccttt ttgtagaatc tacaggtgga	catttagagt gct		343

<210> 143

<211> 354

<212> DNA

<213> Homo sapien

<400> 143

aggtctgatg gcagaaaaac tcagactgtc	tgcaacttta cagatggtgc	attggttcag	60
catcaggagt gggatgggaa ggaaagcaca	ataacaagaa aattgaaaga	tgggaaatta	120
gtggtggagt gtgtcatgaa caatgtcacc	tgtactcgga tctatgaaaa	agtagaataa	180
aaattccatc atcacttttg acaggagtta	attaagagaa tgaccaagct	cagttcaatg	240
agcaaactc catactgttt ctttcttttt	tttttcatta ctgtgttcaa	ttatctttat	300
cataaacatt ttacatgcag ctatttcaaa	gtgtgttgga ttaattagga	tcac	354

<210> 144

<211> 353

<212> DNA

<213> Homo sapien

<400> 144

ggtcaaggac ctgggggacc cccaggtcca	gcagccacat gattctgcag	cagacaggga	60
cctagagcac atctggatct cagccccacc	cctggcaacc tgctgcta	gagaactccc	120
aagatgacag actaagtagg attctgccat	ttagaataat tctggtatcc	tggcggtgc	180
gttaagtgc ttaactttca ttctgtctta	cgatagtctt cagaggtggg	aacagatgaa	240
gaaaccatgc cccagagaag gttaagtgc	ttcctcttta tggagccagt	gttccaacct	300
aggtttgcct gataccagac ctgtggcccc	acctcccatg caggtctctg	tgg	353

<210> 145

<211> 371

<212> DNA

<213> Homo sapien

<400> 145

caggtctgtc ataaaactggt ctggagtctc	tgaagactcc ttgttcacca	aatgcacat	60
ttcctgagac ttgctggcct ctccgttgag	tccacttggc tttctgtcct	ccacagctcc	120
attgccactg ttgatcacta gctttttctt	ctgccacac cttcttcgac	tggtgactgc	180
aatgcaaact gcaagaatca aagccaaggc	caagagggat gccaaagatga	tcagccattc	240

tggaatttgg ggtgtcctta taggaccaga ggttgtgttt gctccacett cttgactccc	300
atgtgagacc tcggccgcga ccacgctaag ccgaattcca gcacactggc ggcccggttac	360
tagtgatcc g	371

<210> 146

<211> 355

<212> DNA

<213> Homo sapien

<400> 146

ggctcctcgt cctcttccca gaggtgtcgg ggcttggccc cagcctccat cttcgtctct	60
caggatggcg agtagcagcg gctccaaggc tgaattcatt gtcggaggga aatataaact	120
ggtacggaag atcgggtctg gctccttcgg ggacatctat ttggcgatca acatcaccaa	180
cggcgaggaa gtggcagtga agctagaatc tcagaaggcc aggcaccccc agttgctgta	240
cgagagcaag ctctataaga ttcttcaagg tgggggtggc atccccca taccgttgta	300
tggtcaggaa aaagactaca atgtactagt catggatctt ctgggaccta gctc	355

<210> 147

<211> 355

<212> DNA

<213> Homo sapien

<400> 147

ggtctgttac aaaatgaaga cagacaacac aacatttact ctgtggagat atcctactca	60
tactatgcac gtgctgtgat tttgaacata actcgtccca aaaacttgtc acgatcatcc	120
tgacttttta ggttggctga tccatcaatc ttgcaactca ctgttacttc tttcccagtg	180
ttgttaggag caaagctgac ctgaacagca accaatggct gtagataccc aacatgcagt	240
tttttcccat aatatgggaa atattttaag tctatcatc cattatgagg ataaactgct	300
acatttggtat tatcttcatt ctttgaaca caatctatcc ttggcactcc ttcag	355

<210> 148

<211> 369

<212> DNA

<213> Homo sapien

<400> 148

aggtctctct cccctctctc ctctcctgcc agccaagtga agacatgctt acttcccctt	60
caccttcctt catgatgtgg gaagagtgtc gcaaccagc cctagccaac accgcatgag	120
agggagtgtg ccgagggtt ctgagaagg tctctccaca tctagaaaga agcgcttaag	180
atgtggcagc ccctcttctt caagtggctc ttgtcctgtt gcctggggag ttctcaaatt	240
gctgcagcag cctccatcca gcctgaggat gacatcaata cacagaggaa gaagagtcag	300
gaaaagatga gagaagttac agactctcct gggcgacccc gagagcttac cattcctcag	360
acttcttca	369

<210> 149

<211> 620

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(620)

<223> n = A,T,C or G

<400> 149

actagtcaaa	aatgctaaaa	taatttgga	gaaaatattt	tttaagtagt	gttatagttt	60
catgtttatc	ttttattatg	ttttgtgaag	ttgtgtcttt	tcactaatta	cctatactat	120
gccaatattt	ccttatatct	atccataaca	tttatactac	atttgaana	naatatgcac	180
gtgaaactta	acactttata	aggtaaaaat	gaggtttcca	anatttaata	atctgatcaa	240
gttcttggtta	tttccaaata	gaatggactt	ggtctgttaa	gggctaagga	gaagaggaag	300
ataaggttaa	aagtgtgtaa	tgaccaaaaca	ttctaaaaga	aatgcaaaaa	aaaagtttat	360
tttcaagcct	tcgaactatt	taaggaaaagc	aaaatcattt	cctaaatgca	tatcatttgt	420
gagaatttct	cattaatatc	ctgaatcatt	catttccacta	aggctcatgt	tnactccgat	480
atgtctctaa	gaaagtacta	tttcatggtc	caaacctggg	tgccatantt	gggtaaaggc	540
tttcccttaa	gtgtgaaant	attttaaagt	aaattttcct	ctttttaaaa	attccttana	600
aggggttaag	gtgttgggga					620

<210> 150

<211> 371

<212> DNA

<213> Homo sapien

<400> 150

ggtcogatca	aaacctgcta	cctccccaag	actttactag	tgccgataaa	ctttctcaaa	60
gagcaaccag	tatcacttcc	ctgtttataa	aaacctatac	catctctttg	ttctttgaac	120
atgctgaaaa	ccacctgggc	tgcatgtatg	cccgaaattg	yaattctttt	ctctcaaatg	180
aaaatttaat	tttagggatt	catttctata	ttttcacata	tgtagtatta	ttatttcctt	240
atatgtgtaa	gggtgaaattt	atgggtattg	agtgtgcaag	aaaatatatt	tttaaagctt	300
tcatttttcc	cccagtgat	gatttagaat	tttttatgta	aataacacaga	atgttttttc	360
ttacttttat	a					371

<210> 151

<211> 4655

<212> DNA

<213> Homo sapien

<400> 151

gggacttgag	ttctgttatt	ttcttaagta	gattcatatt	gtaagggtct	cggggtgggg	60
gggttgga	aatcctggag	ccagaagaaa	ggacagcagc	attgatcaat	cttacagcta	120
acatgttgta	cctggaaaac	aatgcccaga	ctcaatttag	tgagccacag	tacacgaacc	180
tggggtcct	gaacagcatg	gaccagcaga	ttcagaacgg	ctcctcgtec	accagtcctt	240
ataacacaga	ccacgcgcag	aacagcgcta	cggcgccctc	gccctacgca	cagcccagct	300
ccaccttcga	tgctctctct	ccatcacccg	ccatcccttc	caacaccgac	taccagggcc	360
cgcacagttt	cgacgtgtcc	ttccagcagt	cgagcacccg	caagtcggcc	acctggacgt	420
attccactga	actgaagaaa	ctctactgcc	aaattgcaaa	gacatgcccc	atccagatca	480
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agtatgtaga	agatcccac	acaggaagac	agagtgtgct	ggtaccttat	gagccacccc	720
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gagggatgaa	ccgccgtcca	attttaatca	ttgttactct	ggaaaccaga	gatgggcaag	840
tcctgggccc	acgtgtcttt	gaggcccgga	tctgtgcttg	cccaggaaga	gacaggaagg	900
cggatgaaga	tagcatcaga	aagcagcaag	tttcggacag	tacaaagaac	ggtgatggta	960
cgaagcgc	gtttcgtcag	aacacacatg	gtatccagat	gacatccatc	aagaaacgaa	1020
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agtctccatc	ttcatatggg	aacagctccc	cacctctgaa	caaaatgaac	agcatgaaca	1260
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gagacatgaa	tggactcagc	cccacccagg	cactccctcc	cccactctcc	atgccatcca	1440
cctcccactg	cacaccccca	cctccgtatc	ccacagattg	cagcattgtc	agtttcttag	1500
cgagggttgg	ctgttcatca	tgtctggact	atttcacgac	ccaggggctg	accaccatct	1560
atcagattga	gcattactcc	atggatgatc	tggcaagtct	gaaaatccct	gagcaatttc	1620
gacatgcgat	ctggaagggc	atcctggacc	accggcagct	ccacgaattc	tccctcccctt	1680
ctcatctcct	gcggacccca	agcagtgcct	ctacagtcag	tgtgggctcc	agtggagacc	1740
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gcatcaaaga	ggagggggag	tgagcctcac	catgtgagct	cttcctatcc	ctctcctaac	1920
tgccagcccc	ctaaaagcac	tccctgctta	tcttcaaagc	cttctcccta	gctcctcccc	1980
ttcctcttgt	ctgatttctt	aggggaagga	gaagtaagag	gcttacttct	taccctaacc	2040
atctgacctg	gcatctaatt	ctgattctgg	ctttaagcct	tcaaaactat	agcttgcaga	2100
actgtagctt	gccatggcta	ggtagaagtg	agcaaaaaag	agttgggtgt	ctccttaagc	2160
tgcagagatt	tctcattgac	ttttataaag	catgttcacc	cttatagtct	aagactatat	2220
atataaatgt	ataaatatac	agtatagatt	tttgggtggg	gggcattgag	tattgtttaa	2280
aatgtaat	aaatgaaaga	aaattgagtt	gcacttattg	accatttttt	aatttacttg	2340
ttttggatgg	cttgtctata	ctccttccct	taaggggtat	catgtatggg	gataggtatc	2400
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<210> 152
 <211> 586
 <212> PRT
 <213> Homo sapien

<400> 152

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Tyr	Thr	Asn	Leu	Gly	Leu	Leu	Asn	Ser	Met	Asp	Gln	Gln	Ile	Gln	Asn	20	25	30	
Gly	Ser	Ser	Ser	Thr	Ser	Pro	Tyr	Asn	Thr	Asp	His	Ala	Gln	Asn	Ser	35	40	45	
Val	Thr	Ala	Pro	Ser	Pro	Tyr	Ala	Gln	Pro	Ser	Ser	Thr	Phe	Asp	Ala	50	55	60	
Leu	Ser	Pro	Ser	Pro	Ala	Ile	Pro	Ser	Asn	Thr	Asp	Tyr	Pro	Gly	Pro	65	70	75	80
His	Ser	Phe	Asp	Val	Ser	Phe	Gln	Gln	Ser	Ser	Thr	Ala	Lys	Ser	Ala	85	90	95	
Thr	Trp	Thr	Tyr	Ser	Thr	Glu	Leu	Lys	Lys	Leu	Tyr	Cys	Gln	Ile	Ala	100	105	110	
Lys	Thr	Cys	Pro	Ile	Gln	Ile	Lys	Val	Met	Thr	Pro	Pro	Pro	Gln	Gly	115	120	125	
Ala	Val	Ile	Arg	Ala	Met	Pro	Val	Tyr	Lys	Lys	Ala	Glu	His	Val	Thr	130	135	140	
Glu	Val	Val	Lys	Arg	Cys	Pro	Asn	His	Glu	Leu	Ser	Arg	Glu	Phe	Asn	145	150	155	160
Glu	Gly	Gln	Ile	Ala	Pro	Ser	Ser	His	Leu	Ile	Arg	Val	Glu	Gly	Asn	165	170	175	
Ser	His	Ala	Gln	Tyr	Val	Glu	Asp	Pro	Ile	Thr	Gly	Arg	Gln	Ser	Val	180	185	190	
Leu	Val	Pro	Tyr	Glu	Pro	Pro	Gln	Val	Gly	Thr	Glu	Phe	Thr	Thr	Val	195	200	205	
Leu	Tyr	Asn	Phe	Met	Cys	Asn	Ser	Ser	Cys	Val	Gly	Gly	Met	Asn	Arg	210	215	220	
Arg	Pro	Ile	Leu	Ile	Ile	Val	Thr	Leu	Glu	Thr	Arg	Asp	Gly	Gln	Val	225	230	235	240
Leu	Gly	Arg	Arg	Cys	Phe	Glu	Ala	Arg	Ile	Cys	Ala	Cys	Pro	Gly	Arg	245	250	255	
Asp	Arg	Lys	Ala	Asp	Glu	Asp	Ser	Ile	Arg	Lys	Gln	Gln	Val	Ser	Asp	260	265	270	
Ser	Thr	Lys	Asn	Gly	Asp	Gly	Thr	Lys	Arg	Pro	Phe	Arg	Gln	Asn	Thr	275	280	285	
His	Gly	Ile	Gln	Met	Thr	Ser	Ile	Lys	Lys	Arg	Arg	Ser	Pro	Asp	Asp	290	295	300	
Glu	Leu	Val	Tyr	Leu	Pro	Val	Arg	Gly	Arg	Glu	Thr	Tyr	Glu	Met	Leu	305	310	315	320
Val	Lys	Ile	Lys	Glu	Ser	Leu	Glu	Leu	Met	Gln	Tyr	Leu	Leu	Gln	His	325	330	335	
Thr	Ile	Glu	Thr	Tyr	Arg	Gln	Gln	Gln	Gln	Gln	Gln	His	Gln	His	Leu	340	345	350	
Leu	Gln	Lys	Gln	Thr	Ser	Ile	Gln	Ser	Pro	Ser	Ser	Tyr	Gly	Asn	Ser	355	360	365	
Ser	Pro	Pro	Leu	Asn	Lys	Met	Asn	Ser	Met	Asn	Lys	Leu	Pro	Ser	Val	370	375	380	

Ser Gln Leu Ile Asn Pro Gln Gln Arg Asn Ala Leu Thr Pro Thr Thr
 385 390 395 400
 Ile Pro Asp Gly Met Gly Ala Asn Ile Pro Met Met Gly Thr His Met
 405 410 415
 Pro Met Ala Gly Asp Met Asn Gly Leu Ser Pro Thr Gln Ala Leu Pro
 420 425 430
 Pro Pro Leu Ser Met Pro Ser Thr Ser His Cys Thr Pro Pro Pro
 435 440 445
 Tyr Pro Thr Asp Cys Ser Ile Val Ser Phe Leu Ala Arg Leu Gly Cys
 450 455 460
 Ser Ser Cys Leu Asp Tyr Phe Thr Thr Gln Gly Leu Thr Thr Ile Tyr
 465 470 475 480
 Gln Ile Glu His Tyr Ser Met Asp Asp Leu Ala Ser Leu Lys Ile Pro
 485 490 495
 Glu Gln Phe Arg His Ala Ile Trp Lys Gly Ile Leu Asp His Arg Gln
 500 505 510
 Leu His Glu Phe Ser Ser Pro Ser His Leu Leu Arg Thr Pro Ser Ser
 515 520 525
 Ala Ser Thr Val Ser Val Gly Ser Ser Glu Thr Arg Gly Glu Arg Val
 530 535 540
 Ile Asp Ala Val Arg Phe Thr Leu Arg Gln Thr Ile Ser Phe Pro Pro
 545 550 555 560
 Arg Asp Glu Trp Asn Asp Phe Asn Phe Asp Met Asp Ala Arg Arg Asn
 565 570 575
 Lys Gln Gln Arg Ile Lys Glu Glu Gly Glu
 580 585

<210> 153
 <211> 2007
 <212> DNA
 <213> Homo sapien

<400> 153

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tggccagggc	aattttggag	agcaaaaaat	ttgcagttag	agcagtgacc	agggatgtga	180
cttgaccaaa	tgccctggag	ctccagcgcc	ttggagctga	ggtgggtcaa	ggtgacctga	240
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acttctggga	ccctctcaac	caagataagg	aagtgtgtcg	ggggaagctg	gtggcagact	360
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<210> 154

<211> 2148

<212> DNA

<213> Homo sapien

<400> 154

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<210> 155
 <211> 153
 <212> PRT
 <213> Homo sapien

<400> 155
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 20 25 30
 Pro Met Gly Asp Val Pro Met Asp Gly Ile Ser Val Ala Asp Ile Gly
 35 40 45
 Ala Ala Val Ser Ser Ile Phe Asn Ser Pro Glu Glu Phe Leu Gly Lys
 50 55 60
 Ala Val Gly Leu Ser Ala Glu Ala Leu Thr Ile Gln Gln Tyr Ala Asp
 65 70 75 80
 Val Leu Ser Lys Ala Leu Gly Lys Glu Val Arg Asp Ala Lys Ile Thr
 85 90 95
 Pro Glu Ala Phe Glu Lys Leu Gly Phe Pro Ala Ala Lys Glu Ile Ala
 100 105 110
 Asn Met Cys Arg Phe Tyr Glu Met Lys Pro Asp Arg Asp Val Asn Leu
 115 120 125
 Thr His Gln Leu Asn Pro Lys Val Lys Ser Phe Ser Gln Phe Ile Ser
 130 135 140
 Glu Asn Gln Gly Ala Phe Lys Gly Met
 145 150

<210> 156
 <211> 128
 <212> PRT
 <213> Homo sapien

<400> 156
 Met Thr Ser Val Arg Val Ala Ala Tyr Phe Glu Asn Phe Leu Ala Ala
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 Trp Arg Pro Val Lys Ala Ser Asp Gly Asp Tyr Tyr Thr Leu Ala Val
 20 25 30
 Pro Met Gly Asp Val Pro Met Asp Gly Ile Ser Val Ala Asp Ile Gly
 35 40 45
 Ala Ala Val Ser Ser Ile Phe Asn Ser Pro Glu Glu Phe Leu Gly Lys
 50 55 60
 Ala Val Gly Leu Ser Ala Glu Ala Leu Thr Ile Gln Gln Tyr Ala Asp
 65 70 75 80
 Val Leu Ser Lys Ala Leu Gly Lys Glu Val Arg Asp Ala Lys Thr Ile
 85 90 95
 Cys Ala Ile Asp Asp Gln Lys Thr Val Glu Glu Gly Phe Met Glu Asp
 100 105 110
 Val Gly Leu Ser Trp Ser Leu Arg Glu His Asp His Val Ala Gly Ala
 115 120 125

<210> 157
 <211> 424
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (424)
 <223> n = A,T,C or G

<400> 157

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aattcagtc	ccactgttat	attaccttct	ccaggaaccc	tccagtgggg	aaggctgcga	180
tattagattt	ccttgatgac	aaagtgtttg	ttgaaagctg	tgctcagagg	aggtgagagg	240
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tgct						424

<210> 158
 <211> 2099
 <212> DNA
 <213> Homo sapien

<400> 158

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agaagatctg	gctaaacaat	ttctgtatgg	cgaaagaaaa	attctaactt	gtacgccttc	360
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cggaacagtg tggaagcaga aggttttttt aactcatccg tttgccaatc attgcaaaca 2040
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<210> 159

<211> 291

<212> PRT

<213> Homo sapien

<400> 159

Met Asp Trp Gly Thr Leu His Thr Phe Ile Gly Gly Val Asn Lys His
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20 25 30
Val Met Ile Leu Val Val Ala Ala Gln Glu Val Trp Gly Asp Glu Gln
35 40 45
Glu Asp Phe Val Cys Asn Thr Leu Gln Pro Gly Cys Lys Asn Val Cys
50 55 60
Tyr Asp His Phe Phe Pro Val Ser His Ile Arg Leu Trp Ala Leu Gln
65 70 75 80
Leu Ile Phe Val Ser Thr Pro Ala Leu Leu Val Ala Met His Val Ala
85 90 95
Tyr Tyr Arg His Glu Thr Thr Arg Lys Phe Arg Arg Gly Glu Lys Arg
100 105 110
Asn Asp Phe Lys Asp Ile Glu Asp Ile Lys Lys Gln Lys Val Arg Ile
115 120 125
Glu Gly Ser Leu Trp Trp Thr Tyr Thr Ser Ser Ile Phe Phe Arg Ile
130 135 140
Ile Phe Glu Ala Ala Phe Met Tyr Val Phe Tyr Phe Leu Tyr Asn Gly
145 150 155 160
Tyr His Leu Pro Trp Val Leu Lys Cys Gly Ile Asp Pro Cys Pro Asn
165 170 175
Leu Val Asp Cys Phe Ile Ser Arg Pro Thr Glu Lys Thr Val Phe Thr
180 185 190
Ile Phe Met Ile Ser Ala Ser Val Ile Cys Met Leu Leu Asn Val Ala
195 200 205
Glu Leu Cys Tyr Leu Leu Leu Lys Val Cys Phe Arg Arg Ser Lys Arg
210 215 220
Ala Gln Thr Gln Lys Asn His Pro Asn His Ala Leu Lys Glu Ser Lys
225 230 235 240
Gln Asn Glu Met Asn Glu Leu Ile Ser Asp Ser Gly Gln Asn Ala Ile
245 250 255
Thr Gly Ser Gln Ala Lys His Phe Lys Val Lys Cys Ser Cys Val Ile
260 265 270
Arg Arg Leu Leu Ser Ser Pro Glu Gly Asn Thr Asn Leu Lys Val Pro
275 280 285
Ser Val Ala
290

<210> 160

<211> 3951

<212> DNA

<213> Homo sapien

<400> 160

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gaaagagaat	ggaacaaaat	tattataaat	aaatatccaa	agtgctcttc	ttcttagata	2940
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gtattaaaa	gcattgagtt	tttgtacaat	acagataaga	tttttacatg	gtagatcaac	3060
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<210> 161

<211> 943

<212> PRT

<213> Homo sapien

<400> 161

Met Thr Gln Arg Ser Ile Ala Gly Pro Ile Cys Asn Leu Lys Phe Val
 1 5 10 15
 Thr Leu Leu Val Ala Leu Ser Ser Glu Leu Pro Phe Leu Gly Ala Gly
 20 25 30
 Val Gln Leu Gln Asp Asn Gly Tyr Asn Gly Leu Leu Ile Ala Ile Asn
 35 40 45
 Pro Gln Val Pro Glu Asn Gln Asn Leu Ile Ser Asn Ile Lys Glu Met
 50 55 60
 Ile Thr Glu Ala Ser Phe Tyr Leu Phe Asn Ala Thr Lys Arg Arg Val
 65 70 75 80
 Phe Phe Arg Asn Ile Lys Ile Leu Ile Pro Ala Thr Trp Lys Ala Asn
 85 90 95
 Asn Asn Ser Lys Ile Lys Gln Glu Ser Tyr Glu Lys Ala Asn Val Ile
 100 105 110
 Val Thr Asp Trp Tyr Gly Ala His Gly Asp Asp Pro Tyr Thr Leu Gln
 115 120 125
 Tyr Arg Gly Cys Gly Lys Glu Gly Lys Tyr Ile His Phe Thr Pro Asn
 130 135 140
 Phe Leu Leu Asn Asp Asn Leu Thr Ala Gly Tyr Gly Ser Arg Gly Arg
 145 150 155 160
 Val Phe Val His Glu Trp Ala His Leu Arg Trp Gly Val Phe Asp Glu
 165 170 175
 Tyr Asn Asn Asp Lys Pro Phe Tyr Ile Asn Gly Gln Asn Gln Ile Lys
 180 185 190
 Val Thr Arg Cys Ser Ser Asp Ile Thr Gly Ile Phe Val Cys Glu Lys
 195 200 205
 Gly Pro Cys Pro Gln Glu Asn Cys Ile Ile Ser Lys Leu Phe Lys Glu
 210 215 220
 Gly Cys Thr Phe Ile Tyr Asn Ser Thr Gln Asn Ala Thr Ala Ser Ile
 225 230 235 240
 Met Phe Met Gln Ser Leu Ser Ser Val Val Glu Phe Cys Asn Ala Ser
 245 250 255
 Thr His Asn Gln Glu Ala Pro Asn Leu Gln Asn Gln Met Cys Ser Leu
 260 265 270
 Arg Ser Ala Trp Asp Val Ile Thr Asp Ser Ala Asp Phe His His Ser
 275 280 285
 Phe Pro Met Asn Gly Thr Glu Leu Pro Pro Pro Thr Phe Ser Leu
 290 295 300

Val Glu Ala Gly Asp Lys Val Val Cys Leu Val Leu Asp Val Ser Ser
 305 310 315 320
 Lys Met Ala Glu Ala Asp Arg Leu Leu Gln Leu Gln Gln Ala Ala Glu
 325 330 335
 Phe Tyr Leu Met Gln Ile Val Glu Ile His Thr Phe Val Gly Ile Ala
 340 345 350
 Ser Phe Asp Ser Lys Gly Glu Ile Arg Ala Gln Leu His Gln Ile Asn
 355 360 365
 Ser Asn Asp Asp Arg Lys Leu Leu Val Ser Tyr Leu Pro Thr Thr Val
 370 375 380
 Ser Ala Lys Thr Asp Ile Ser Ile Cys Ser Gly Leu Lys Lys Gly Phe
 385 390 395 400
 Glu Val Val Glu Lys Leu Asn Gly Lys Ala Tyr Gly Ser Val Met Ile
 405 410 415
 Leu Val Thr Ser Gly Asp Asp Lys Leu Leu Gly Asn Cys Leu Pro Thr
 420 425 430
 Val Leu Ser Ser Gly Ser Thr Ile His Ser Ile Ala Leu Gly Ser Ser
 435 440 445
 Ala Ala Pro Asn Leu Glu Glu Leu Ser Arg Leu Thr Gly Gly Leu Lys
 450 455 460
 Phe Phe Val Pro Asp Ile Ser Asn Ser Asn Ser Met Ile Asp Ala Phe
 465 470 475 480
 Ser Arg Ile Ser Ser Gly Thr Gly Asp Ile Phe Gln Gln His Ile Gln
 485 490 495
 Leu Glu Ser Thr Gly Glu Asn Val Lys Pro His His Gln Leu Lys Asn
 500 505 510
 Thr Val Thr Val Asp Asn Thr Val Gly Asn Asp Thr Met Phe Leu Val
 515 520 525
 Thr Trp Gln Ala Ser Gly Pro Pro Glu Ile Ile Leu Phe Asp Pro Asp
 530 535 540
 Gly Arg Lys Tyr Tyr Thr Asn Asn Phe Ile Thr Asn Leu Thr Phe Arg
 545 550 555 560
 Thr Ala Ser Leu Trp Ile Pro Gly Thr Ala Lys Pro Gly His Trp Thr
 565 570 575
 Tyr Thr Leu Asn Asn Thr His His Ser Leu Gln Ala Leu Lys Val Thr
 580 585 590
 Val Thr Ser Arg Ala Ser Asn Ser Ala Val Pro Pro Ala Thr Val Glu
 595 600 605
 Ala Phe Val Glu Arg Asp Ser Leu His Phe Pro His Pro Val Met Ile
 610 615 620
 Tyr Ala Asn Val Lys Gln Gly Phe Tyr Pro Ile Leu Asn Ala Thr Val
 625 630 635 640
 Thr Ala Thr Val Glu Pro Glu Thr Gly Asp Pro Val Thr Leu Arg Leu
 645 650 655
 Leu Asp Asp Gly Ala Gly Ala Asp Val Ile Lys Asn Asp Gly Ile Tyr
 660 665 670
 Ser Arg Tyr Phe Phe Ser Phe Ala Asn Gly Arg Tyr Ser Leu Lys
 675 680 685
 Val His Val Asn His Ser Pro Ser Ile Ser Thr Pro Ala His Ser Ile
 690 695 700
 Pro Gly Ser His Ala Met Tyr Val Pro Gly Tyr Thr Ala Asn Gly Asn
 705 710 715 720
 Ile Gln Met Asn Ala Pro Arg Lys Ser Val Gly Arg Asn Glu Glu Glu
 725 730 735
 Arg Lys Trp Gly Phe Ser Arg Val Ser Ser Gly Gly Ser Phe Ser Val

	740		745		750
Leu Gly Val	Pro Ala Gly	Pro His	Pro Asp Val	Phe Pro	Pro Cys Lys
755		760		765	
Ile Ile Asp	Leu Glu Ala	Val Lys	Val Glu Glu	Glu Leu Thr	Leu Ser
770		775		780	
Trp Thr Ala	Pro Gly Glu	Asp Phe	Asp Gln Gly	Gln Ala Thr	Ser Tyr
785		790		795	800
Glu Ile Arg	Met Ser Lys	Ser Leu	Gln Asn Ile	Gln Asp Asp	Phe Asn
	805		810		815
Asn Ala Ile	Leu Val Asn	Thr Ser	Lys Arg Asn	Pro Gln Gln	Ala Gly
	820		825		830
Ile Arg Glu	Ile Phe Thr	Phe Ser	Pro Gln Ile	Ser Thr Asn	Gly Pro
	835		840		845
Glu His Gln	Pro Asn Gly	Glu Thr	His Glu Ser	His Arg Ile	Tyr Val
	850		855		860
Ala Ile Arg	Ala Met Asp	Arg Asn	Ser Leu Gln	Ser Ala Val	Ser Asn
	865		870		875
Ile Ala Gln	Ala Pro Leu	Phe Ile	Pro Pro Asn	Ser Asp Pro	Val Pro
	885		890		895
Ala Arg Asp	Tyr Leu Ile	Leu Lys	Gly Val Leu	Thr Ala Met	Gly Leu
	900		905		910
Ile Gly Ile	Ile Cys Leu	Ile Ile	Val Val Thr	His His Thr	Leu Ser
	915		920		925
Arg Lys Lys	Arg Ala Asp	Lys Lys	Glu Asn Gly	Thr Lys Leu	Leu
	930		935		940

<210> 162
 <211> 498
 <212> DNA
 <213> Homo sapien

<400> 162

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agcccctcaa	gtcgggtatg	aaggagctgg	cgtgttccg	ggagaaggtc	actgagcagc	120
accggcagat	gggcaagggt	ggcaagcatc	accttggcct	ggaggagccc	aagaagctgc	180
gaccaccccc	tgccaggact	ccctgccaac	aggaactgga	ccaggtcctg	gagcggatct	240
ccaccatgcg	ccttccggat	gagcggggcc	ctctggagca	cctctactcc	ctgcacatcc	300
ccaactgtga	caagcatggc	ctgtacaacc	tcaaacagtg	gcaagatgtc	tctgaacggg	360
cagcgtgggg	agtgtgtgtg	tgtgaacccc	aacacgggga	agctgatcca	gggagccccc	420
accatccggg	gggaccccga	gtgtcatctc	ttctacaatg	agcagcagga	ggctcggggg	480
gtgcacaccc	cagcggat					498

<210> 163
 <211> 1128
 <212> DNA
 <213> Homo sapien

<400> 163

gccacctggc	cctcctgatc	gacgacacac	gcacttgaaa	cttgttctca	gggtgtgtgg	60
aatcaacttt	ccggaagcaa	ccagcccacc	agaggaggtc	ccgagcgcca	gaggagacga	120
tgcagcggag	actggttcag	cagtggagcg	tgcgggtgtt	cctgctgagc	tacgcgggtgc	180
cctcctgcgg	gcgtcgggtg	gaggggtctca	gcgcgcgcct	caaaagagct	gtgtctgaac	240
atcagctcct	ccatgacaag	gggaagtcca	tccaagattt	acggcgacga	ttcttccttc	300
accatctgat	cgcagaaatc	cacacagctg	aaatcagagc	tacctcggag	gtgtccccta	360
actccaagcc	ctctcccaac	acaaagaacc	accccgctcg	atttgggtct	gatgatgagg	420

gcagatacct	aactcaggaa	actaacaagg	tggagacgta	caaagagcag	ccgctcaaga	480
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aattattatt	atcacattta	ccataattta	ttttgtccat	tgatgtattt	attttgtaaa	900
tgtatcttgg	tgctgctgaa	tttctatatt	ttttgtaaca	taatgcactt	tagatataca	960
tatcaagtat	gttgataaat	gacacaatga	agtgtctcta	ttttgtgggt	gatttttaag	1020
aatgcctaaa	tataattatc	caaattgatt	ttcctttgtg	catgtaaaaa	taacagtatt	1080
ttaaatttgt	aaagaatgtc	taataaaata	taatctaatt	acatcatg		1128

<210> 164

<211> 1310

<212> DNA

<213> Homo sapien

<400> 164

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ttagccctgt	tccacgaacc	caggagaact	gctggccaga	ttaattagac	attgctatgg	120
gagacgtgta	aacacactac	ttatcattga	tgcatatata	aaaccatttt	attttcgcta	180
ttatttcaga	ggaagcgctt	ctgatttgtt	tcttttttcc	ctttttgctc	tttctggctg	240
tgtggtttgg	agaaagcaca	gttgagtag	cgggttgcta	aataagtccc	gagcgcgagc	300
ggagacgatg	cagcggagac	tggttcagca	gtggagcgtc	gcggtgttcc	tgctgagcta	360
cgcggtgccc	tcctgcgggc	gctcgttgga	gggtctcagc	cgccgcctca	aaagagctgt	420
gtctgaacat	cagtcctcc	atgacaagg	gaagtccatc	caagatttac	ggcgacgatt	480
cttccttcac	catctgatcg	cagaaatcca	cacagctgaa	atcagagcta	cctcggaggt	540
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gaaagtatta	gaaatattta	ttgtctgtaa	atactgtaaa	tgatttgga	taaaactgtc	960
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ttttaatgaa	tgccataata	taattatcca	aattgatttt	cctttgtgcc	cgtaaaaata	1260
acagtatttt	aaatttgtaa	agaatgtcta	ataaaatata	atctaattac		1310

<210> 165

<211> 177

<212> PRT

<213> Homo sapien

<400> 165

Met	Gln	Arg	Arg	Leu	Val	Gln	Gln	Trp	Ser	Val	Ala	Val	Phe	Leu	Leu
1				5				10					15		
Ser	Tyr	Ala	Val	Pro	Ser	Cys	Gly	Arg	Ser	Val	Glu	Gly	Leu	Ser	Arg
			20					25					30		
Arg	Leu	Lys	Arg	Ala	Val	Ser	Glu	His	Gln	Leu	Leu	His	Asp	Lys	Gly
			35					40					45		
Lys	Ser	Ile	Gln	Asp	Leu	Arg	Arg	Arg	Phe	Phe	Leu	His	His	Leu	Ile

50	55	60
Ala Glu Ile His Thr	Ala Glu Ile Arg	Ala Thr Ser Glu Val Ser Pro
65	70	75
Asn Ser Lys Pro Ser	Pro Asn Thr Lys	Asn His Pro Val Arg Phe Gly
85	90	95
Ser Asp Asp Glu Gly Arg	Tyr Leu Thr Gln Glu Thr Asn Lys Val Glu	
100	105	110
Thr Tyr Lys Glu Gln Pro	Leu Lys Thr Pro Gly Lys Lys Lys Gly	
115	120	125
Lys Pro Gly Lys Arg Lys	Glu Gln Glu Lys Lys Lys Arg Arg Thr Arg	
130	135	140
Ser Ala Trp Leu Asp Ser	Gly Val Thr Gly Ser Gly Leu Glu Gly Asp	
145	150	155
His Leu Ser Asp Thr	Ser Thr Thr Ser Leu Glu Leu Asp Ser Arg Arg	
165	170	175

His

<210> 166
 <211> 177
 <212> PRT
 <213> Homo sapien

<400> 166
Met Gln Arg Arg Leu Val Gln Gln Trp Ser Val Ala Val Phe Leu Leu
1 5 10 15
Ser Tyr Ala Val Pro Ser Cys Gly Arg Ser Val Glu Gly Leu Ser Arg
20 25 30
Arg Leu Lys Arg Ala Val Ser Glu His Gln Leu Leu His Asp Lys Gly
35 40 45
Lys Ser Ile Gln Asp Leu Arg Arg Arg Phe Phe Leu His His Leu Ile
50 55 60
Ala Glu Ile His Thr Ala Glu Ile Arg Ala Thr Ser Glu Val Ser Pro
65 70 75 80
Asn Ser Lys Pro Ser Pro Asn Thr Lys Asn His Pro Val Arg Phe Gly
85 90 95
Ser Asp Asp Glu Gly Arg Tyr Leu Thr Gln Glu Thr Asn Lys Val Glu
100 105 110
Thr Tyr Lys Glu Gln Pro Leu Lys Thr Pro Gly Lys Lys Lys Gly
115 120 125
Lys Pro Gly Lys Arg Lys Glu Gln Glu Lys Lys Lys Arg Arg Thr Arg
130 135 140
Ser Ala Trp Leu Asp Ser Gly Val Thr Gly Ser Gly Leu Glu Gly Asp
145 150 155 160
His Leu Ser Asp Thr Ser Thr Thr Ser Leu Glu Leu Asp Ser Arg Arg
165 170 175

His

<210> 167
 <211> 3362
 <212> DNA
 <213> Homo sapien

<400> 167

cacaatgtat	gcagcaggct	cagtgtgagt	gaactggagg	cttctctaca	acatgacca	60
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<210> 168

<211> 2784

<212> DNA

<213> Homo sapien

<400> 168

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2784

<210> 169

<211> 592

<212> PRT

<213> Homo sapien

<400> 169

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 35 40 45
 Pro Gln Val Pro Glu Asn Gln Asn Leu Ile Ser Asn Ile Lys Glu Met
 50 55 60
 Ile Thr Glu Ala Ser Phe Tyr Leu Phe Asn Ala Thr Lys Arg Arg Val
 65 70 75 80
 Phe Phe Arg Asn Ile Lys Ile Leu Ile Pro Ala Thr Trp Lys Ala Asn
 85 90 95
 Asn Asn Ser Lys Ile Lys Gln Glu Ser Tyr Glu Lys Ala Asn Val Ile
 100 105 110
 Val Thr Asp Trp Tyr Gly Ala His Gly Asp Asp Pro Tyr Thr Leu Gln
 115 120 125
 Tyr Arg Gly Cys Gly Lys Glu Gly Lys Tyr Ile His Phe Thr Pro Asn
 130 135 140
 Phe Leu Leu Asn Asp Asn Leu Thr Ala Gly Tyr Gly Ser Arg Gly Arg
 145 150 155 160
 Val Phe Val His Glu Trp Ala His Leu Arg Trp Gly Val Phe Asp Glu
 165 170 175
 Tyr Asn Asn Asp Lys Pro Phe Tyr Ile Asn Gly Gln Asn Gln Ile Lys
 180 185 190
 Val Thr Arg Cys Ser Ser Asp Ile Thr Gly Ile Phe Val Cys Glu Lys
 195 200 205
 Gly Pro Cys Pro Gln Glu Asn Cys Ile Ile Ser Lys Leu Phe Lys Glu
 210 215 220
 Gly Cys Thr Phe Ile Tyr Asn Ser Thr Gln Asn Ala Thr Ala Ser Ile
 225 230 235 240
 Met Phe Met Gln Ser Leu Ser Ser Val Val Glu Phe Cys Asn Ala Ser
 245 250 255
 Thr His Asn Gln Glu Ala Pro Asn Leu Gln Asn Gln Met Cys Ser Leu
 260 265 270
 Arg Ser Ala Trp Asp Val Ile Thr Asp Ser Ala Asp Phe His His Ser
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 Phe Pro Met Asn Gly Thr Glu Leu Pro Pro Pro Pro Thr Phe Ser Leu
 290 295 300
 Val Glu Ala Gly Asp Lys Val Val Cys Leu Val Leu Asp Val Ser Ser
 305 310 315 320
 Lys Met Ala Glu Ala Asp Arg Leu Leu Gln Leu Gln Gln Ala Ala Glu
 325 330 335
 Phe Tyr Leu Met Gln Ile Val Glu Ile His Thr Phe Val Gly Ile Ala
 340 345 350
 Ser Phe Asp Ser Lys Gly Glu Ile Arg Ala Gln Leu His Gln Ile Asn
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 Ser Asn Asp Asp Arg Lys Leu Leu Val Ser Tyr Leu Pro Thr Thr Val

370	375	380
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Leu Val Thr Ser Gly Asp Asp Lys Leu Gly Asn Cys Leu Pro Thr		415
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Val Leu Ser Ser Gly Ser Thr Ile His Ser Ile Ala Leu Gly Ser Ser		430
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Ala Ala Pro Asn Leu Glu Glu Leu Ser Arg Leu Thr Gly Gly Leu Lys		445
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Phe Phe Val Pro Asp Ile Ser Asn Ser Asn Ser Met Ile Asp Ala Phe		460
465	470	475
Ser Arg Ile Ser Ser Gly Thr Gly Asp Ile Phe Gln Gln His Ile Gln		480
	485	490
Leu Glu Ser Thr Gly Glu Asn Val Lys Pro His His Gln Leu Lys Asn		495
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Thr Val Thr Val Asp Asn Thr Val Gly Asn Asp Thr Met Phe Leu Val		510
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Gly Arg Lys Tyr Tyr Thr Asn Asn Phe Ile Thr Asn Leu Thr Phe Arg		540
545	550	555
Thr Ala Ser Leu Trp Ile Pro Gly Thr Ala Lys Pro Gly His Trp Thr		560
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<210> 170

<211> 791

<212> PRT

<213> Homo sapien

<400> 170

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Pro Gln Val Pro Glu Asn Gln Asn Leu Ile Ser Asn Ile Lys Glu Met	40
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Ile Thr Glu Ala Ser Phe Tyr Leu Phe Asn Ala Thr Lys Arg Arg Val	55
65	70
Phe Phe Arg Asn Ile Lys Ile Leu Ile Pro Ala Thr Trp Lys Ala Asn	75
	85
Asn Asn Ser Lys Ile Lys Gln Glu Ser Tyr Glu Lys Ala Asn Val Ile	90
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Val Thr Asp Trp Tyr Gly Ala His Gly Asp Asp Pro Tyr Thr Leu Gln	105
	115
Tyr Arg Gly Cys Gly Lys Glu Gly Lys Tyr Ile His Phe Thr Pro Asn	120
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Phe Leu Leu Asn Asp Asn Leu Thr Ala Gly Tyr Gly Ser Arg Gly Arg	135
145	150
Val Phe Val His Glu Trp Ala His Leu Arg Trp Gly Val Phe Asp Glu	155
	160

Ala Phe Val Glu Arg Asp Ser Leu His Phe Pro His Pro Val Met Ile
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 Tyr Ala Asn Val Lys Gln Gly Phe Tyr Pro Ile Leu Asn Ala Thr Val
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 Thr Ala Thr Val Glu Pro Glu Thr Gly Asp Pro Val Thr Leu Arg Leu
 645 650 655
 Leu Asp Asp Gly Ala Gly Ala Asp Val Ile Lys Asn Asp Gly Ile Tyr
 660 665 670
 Ser Arg Tyr Phe Phe Ser Phe Ala Ala Asn Gly Arg Tyr Ser Leu Lys
 675 680 685
 Val His Val Asn His Ser Pro Ser Ile Ser Thr Pro Ala His Ser Ile
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 Pro Gly Ser His Ala Met Tyr Val Pro Gly Tyr Thr Ala Asn Gly Asn
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 Ile Gln Met Asn Ala Pro Arg Lys Ser Val Gly Arg Asn Glu Glu Glu
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 Arg Lys Trp Gly Phe Ser Arg Val Ser Ser Gly Gly Ser Phe Ser Val
 740 745 750
 Leu Gly Val Pro Ala Gly Pro His Pro Asp Val Phe Pro Pro Cys Lys
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<210> 171

<211> 1491

<212> DNA

<213> Homo sapien

<400> 171

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1491

<210> 172

<211> 364

<212> PRT

<213> Homo sapien

<400> 172

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Leu	Ala	Ile	Glu	Ala	Gly	Phe	His	His	Ile	Asp	Ser	Ala	His	Val	Tyr
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Asn	Asn	Glu	Glu	Gln	Val	Gly	Leu	Ala	Ile	Arg	Ser	Lys	Ile	Ala	Asp
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Gly	Ser	Val	Lys	Arg	Glu	Asp	Ile	Phe	Tyr	Thr	Ser	Lys	Leu	Trp	Ser
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Asn	Ser	His	Arg	Pro	Glu	Leu	Val	Arg	Pro	Ala	Leu	Glu	Arg	Ser	Leu
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Val	Ser	Val	Lys	Pro	Gly	Glu	Glu	Val	Ile	Pro	Lys	Asp	Glu	Asn	Gly
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Lys	Ile	Leu	Phe	Asp	Thr	Val	Asp	Leu	Cys	Ala	Thr	Trp	Glu	Ala	Met
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Tyr	Ser	Ala	Leu	Gly	Ser	His	Arg	Glu	Glu	Pro	Trp	Val	Asp	Pro	Asn
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Ser	Pro	Val	Leu	Leu	Glu	Asp	Pro	Val	Leu	Cys	Ala	Leu	Ala	Lys	Lys
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His	Lys	Arg	Thr	Pro	Ala	Leu	Ile	Ala	Leu	Arg	Tyr	Gln	Leu	Gln	Arg
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Ile	Asp	Gly	Leu	Asn	Arg	Asn	Val	Arg	Tyr	Leu	Thr	Leu	Asp	Ile	Phe
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Leu Arg Ser Ala Pro Leu Gly Pro Ala Pro Pro Val Asn Met Ile Arg

35	40	45
Cys Gly Leu Ala Cys Glu Arg Cys Arg Trp Ile Leu Pro Leu Leu Leu		
50	55	60
Leu Ser Ala Ile Ala Phe Asp Ile Ile Ala Leu Ala Gly Arg Gly Trp		
65	70	75 80
Leu Gln Ser Ser Asp His Gly Gln Thr Ser Ser Leu Trp Trp Lys Cys		
	85 90	95
Ser Gln Glu Gly Gly Gly Ser Gly Ser Tyr Glu Glu Gly Cys Gln Ser		
100	105	110
Leu Met Glu Tyr Ala Trp Gly Arg Ala Ala Ala Met Leu Phe Cys		
115	120	125
Gly Phe Ile Ile Leu Val Ile Cys Phe Ile Leu Ser Phe Phe Ala Leu		
130	135	140
Cys Gly Pro Gln Met Leu Val Phe Leu Arg Val Ile Gly Gly Leu Leu		
145	150	155 160
Ala Leu Ala Ala Val Phe Gln Ile Ile Ser Leu Val Ile Tyr Pro Val		
165	170	175
Lys Tyr Thr Gln Thr Phe Thr Leu His Ala Asn Pro Ala Val Thr Tyr		
180	185	190
Ile Tyr Asn Trp Ala Tyr Gly Phe Gly Trp Ala Ala Thr Ile Ile Leu		
195	200	205
Ile Gly Cys Ala Phe Phe Phe Cys Cys Leu Pro Asn Tyr Glu Asp Asp		
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<400> 175

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<210> 176
<211> 580
<212> PRT
<213> Homo sapiens

<400> 176

Glu Lys Ser Ile Thr Ile Leu Ser Thr Pro Glu Gly Thr Ser Ala Ala
245 250 255

Cys Lys Ser Ile Leu Glu Ile Met His Lys Glu Ala Gln Asp Ile Lys
 260 265 270
 Phe Thr Glu Glu Ile Pro Leu Lys Ile Leu Ala His Asn Asn Phe Val
 275 280 285
 Gly Arg Leu Ile Gly Lys Glu Gly Arg Asn Leu Lys Lys Ile Glu Gln
 290 295 300
 Asp Thr Asp Thr Lys Ile Thr Ile Ser Pro Leu Gln Glu Leu Thr Leu
 305 310 315 320
 Tyr Asn Pro Glu Arg Thr Ile Thr Val Lys Gly Asn Val Glu Thr Cys
 325 330 335
 Ala Lys Ala Glu Glu Glu Ile Met Lys Lys Ile Arg Glu Ser Tyr Glu
 340 345 350
 Asn Asp Ile Ala Ser Met Asn Leu Gln Ala His Leu Ile Pro Gly Leu
 355 360 365
 Asn Leu Asn Ala Leu Gly Leu Phe Pro Pro Thr Ser Gly Met Pro Pro
 370 375 380
 Pro Thr Ser Gly Pro Pro Ser Ala Met Thr Pro Pro Tyr Pro Gln Phe
 385 390 395 400
 Glu Gln Ser Glu Thr Glu Thr Val His Gln Phe Ile Pro Ala Leu Ser
 405 410 415
 Val Gly Ala Ile Ile Gly Lys Gln Gly Gln His Ile Lys Gln Leu Ser
 420 425 430
 Arg Phe Ala Gly Ala Ser Ile Lys Ile Ala Pro Ala Glu Ala Pro Asp
 435 440 445
 Ala Lys Val Arg Met Val Ile Ile Thr Gly Pro Pro Glu Ala Gln Phe
 450 455 460
 Lys Ala Gln Gly Arg Ile Tyr Gly Lys Ile Lys Glu Glu Asn Phe Val
 465 470 475 480
 Ser Pro Lys Glu Glu Val Lys Leu Glu Ala His Ile Arg Val Pro Ser
 485 490 495
 Phe Ala Ala Gly Arg Val Ile Gly Lys Gly Gly Lys Thr Val Asn Glu
 500 505 510
 Leu Gln Asn Leu Ser Ser Ala Glu Val Val Val Pro Arg Asp Gln Thr
 515 520 525
 Pro Asp Glu Asn Asp Gln Val Val Val Lys Ile Thr Gly His Phe Tyr
 530 535 540

Ala Cys Gln Val Ala Gln Arg Lys Ile Gln Glu Ile Leu Thr Gln Val
 545 550 555 560

Lys Gln His Gln Gln Gln Lys Ala Leu Gln Ser Gly Pro Pro Gln Ser
 565 570 575

Arg Arg Lys

<210> 177

<211> 401

<212> DNA

<213> Homo sapiens

<400> 177

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 cacacagcaa aaaattgttt actttgttgg acaaaccaaa tcagttctca aaaaatgacc 180
 ggtgcttata aaaagttata aatatcgagt agctctaaaa caaaccacct gaccaagagg 240
 gaagtgagct tgtgcttagt atttacattg gatgccagtt ttgtaatcac tgacttatgt 300
 gcaaactggg gcagaaattc tataaactct ttgctgtttt tgatacctgc tttttgtttc 360
 attttggttt gttttgtaaa aatgataaaa cttcagaaaa t 401

<210> 178

<211> 561

<212> DNA

<213> Homo sapiens

<400> 178

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 gcccgctatg ggacaggggt ctttgccag aatgagtacc tacgctatca ggaggccctg 120
 agtgaagctgg cactgcgggt taaagcacga attgggagct ctacgcgaca tcaccagtca 180
 gcagccaaag acctaactca gtcccctgag gtctcccca caaccatcca ggtgacatac 240
 ctcccctcca gtcagaagag taaacgtgcc aagcacttcc ttgaattgaa gagctttaag 300
 gataactata acacattgga gagtactctg tgacggagct gaaggactct tgccgtagat 360
 taagccagtc agttgcaatg tgcaagacag gctgcttgcc gggccgcct cggaacatct 420
 ggcccagcag gccagactg tatccatcca agttcccggt gtatccagag ttcttagagc 480
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 gactattttc cccagtagc g 561

<210> 179

<211> 521

<212> DNA

<213> Homo sapiens

<400> 179

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 gatcgagcaa tggcttcagg acatgggttc tcttctcctg tgatcattca agtgctcact 120
 gcatgaagac tggcttgtct cagtgtttca acctcaccag ggctgtctct tgggtccacac 180
 ctgcctccct gttagtgcg tatgacagcc cccatcaaata gaccttggcc aagtcacggg 240
 ttctctgtgg tcaaggttgg ttggctgatt ggtggaaagt aggggtggacc aaaggaggcc 300
 acgtgagcag tcagcaccag ttctgcacca gcagcgctc cgtcctagtg ggtgttctg 360
 tttctcctgg ccctgggtgg gctagggcct gattcgggaa gatgccttg cagggagggg 420
 aggataagtg ggatctacca attgattctg gcaaaacaat ttctaagatt tttttgcttt 480

atgtgggaaa cagatctaaa tctcatttta tgctgtattt t

521

<210> 180

<211> 417

<212> DNA

<213> Homo sapiens

<400> 180

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tcctgggccg cctggcggcc atcgtggcta aacagggtact gctggggcgg aagggtggtgg 120
tcgtacgctg tgaaggcatc aacatttctg gcaatttcta cagaaacaag ttgaagtacc 180
tggttttctc cgcgaagcgg atgaacacca acccttcccg aggccctac cacttccggg 240
ccccagccg catcttctg cggaccgtgc gaggtatgct gcccacaaa accaagcgag 300
gccaggccgc tctggaccgt ctcaagggtg ttgacggcat cccaccgcc tacgacaaga 360
aaaagcggat ggtggttcct gctgccctca aggtcgtgcg tctgaagcct acaagaa 417

<210> 181

<211> 283

<212> DNA

<213> Homo sapiens

<220>

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<222> (35)

<223> n=A,T,C or G

<400> 181

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caagaactca agtgtaactg tgataaaata acctttccca ggtatatggt caggatggtg 120
tgtaattctca gaatacacag gtgacataga tatgatatga caactggtaa tgggtggattc 180
atttacattg tttaacttc tatgaccagg ccttaaggga aggtcagttt tttaaaaaac 240
caagtagtgt cttcctacct atctccagat acatgtcaaa aaa 283

<210> 182

<211> 401

<212> DNA

<213> Homo sapiens

<400> 182

atattcttgc tgcttatgca gctgacattg ttgccctccc taaagcaacc aagtagcctt 60
tatttcccac agtgaaagaa aacgctggcc tatcagttac attacaaaag gcagatttca 120
agaggattga gtaagtagtt ggatggcttt cataaaaaa agaattcaag aagaggattc 180
atgctttaag aaacatttgt tataattcc tcacaaatta tacctgggat aaaaactatg 240
tagcaggcag tgtgttttcc ttccatgtct ctctgcacta cctgcagtgt gtcctctgag 300
gctgcaagtc tgtctatct gaattcccag cagaagcact aagaagctcc accctatcac 360
ctagcagata aaactatggg gaaaacttaa atctgtgcat a 401

<210> 183

<211> 366

<212> DNA

<213> Homo sapiens

<220>

<221> unsure

<222> (325)

<223> n=A,T,C or G

<400> 183

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accatcatgc tttgatgttc ccctgtcttt ctctctcttg ctctcaagag caaagggttaa 120
tttaaggaca aagatgaagt cactgtaaac taatctgtca ttgtttttac ctctcttttc 180
tttttcagtg cagaaattaa aagtaagtat aaagcacctg gattgggagt gtttttgcgt 240
gtgtcggaat cactggtaaa tgttggtga gaacaatccc tccccttgca cttgtgaaaa 300
cactttgagc gctttaagag attancctga gaaataatta aatatctttt ctcttcaaaa 360
aaaaaa 366
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<210> 184

<211> 370

<212> DNA

<213> Homo sapiens

<400> 184

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tttaataatt gtactgagag aaactgctta cgtacacatt gcagatcaaa tatttgaggt 120
taaaatgtta gtctacatag atgggtgatt gtaactttat tgccattaaa agatttcaaa 180
ttgcattcat gtttctgtgt acacataatg aaaaatgggc aaataatgaa gatctctcct 240
tcagtctgct ctgtttaatt ctgctgtctg ctcttctcta atgctgcgtc cctaattgta 300
cacagtttag tgatatctag gagtataaag ttgtcgccca tcaataaaaa tcacaaagtt 360
ggtttaaaaa 370
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<210> 185

<211> 107

<212> DNA

<213> Homo sapiens

<400> 185

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ctcatattat tttccttttg agaaattgga aactctttct gttgctatta tattaataaa 60
gttggtgttt attttctggt agtcaccttc cccatttaaa aaaaaaa 107
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<210> 186

<211> 309

<212> DNA

<213> Homo sapiens

<400> 186

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agagggccac aggggtggcc gggagtgtgc agctgatgcc tgctgagagg caggaattgt 120
gccagtgaat gacagtcagt agggagtgtc tcttcttggg gaggaagaa ggtagagcct 180
ttctgtctga atgaaaggcc aaggctacag tacagggccc cgcccagcc aggggtgttaa 240
tgcccacgta gtggaggcct ctggcagatc ctgcattcca aggtcactgg actgtacgtt 300
tttatggtt 309
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<210> 187

<211> 477

<212> DNA

<213> Homo sapiens

<400> 187

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tccaacctcg ggccagtgtc ttcaggcttt actggggacc tgcgagctgg cctaattgtg 120
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```

tggcctgcaa gccaggccat ccctgggcgc cacagacgag ctccgagcca ggtcaggctt 180
cggaggccac aagctcagcc tcaggcccag gcaactgattg tggcagaggg gccactaccc 240
aaggtctagc taggcccacg acctagttac ccagacagtg agaagcccct ggaaggcaga 300
aaagtgggga gcatggcaga caggggaaggg aaacattttc agggaaaaga catgtatcac 360
atgtcttcag aagcaagtca ggtttcatgt aaccgagtgt cctcttgctg gtccaaaagt 420
agcccagggc ttagcacacg gttcacagt gattttgtgt tcagccgtga gtcacac 477

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<210> 188

<211> 220

<212> DNA

<213> Homo sapiens

<400> 188

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ttaaataagt accctgtgag tatgagataa attagtgaca atcagaacaa gtttcagtat 120
cagatgttca agaggaagtt gctattgcat tgattttaat attgtacat aaacactgat 180
ttttttgagc attattttgt attgtgtgta ctttaatacc 220

```

<210> 189

<211> 417

<212> DNA

<213> Homo sapiens

<220>

<221> unsure

<222> (76)

<223> n=A,T,C or G

<221> unsure

<222> (77)

<223> n=A,T,C or G

<400> 189

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tatcattatt ctagtccctt gaatttgtaa ggggaaaaaa aacaaaaaça aaaacttacg 180
atgcaacttt ctccagcaca tcagatttca aattgaaaat taaagacatg ctatggtaat 240
gcacttgcta gtactacaca ctttgtacaa caaaaaacag aggcaagaaa caacggaaag 300
agaaaagcct tcctttgttg gcccttaaac tgagtcaaga tctgaaatgt agagatgatc 360
tctgacgata cctgtatgtt cttattgtgt aaataaaatt gctggtatga aatgaca 417

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<210> 190

<211> 497

<212> DNA

<213> Homo sapiens

<400> 190

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aacgcaggag ctgtcattga ctggcccaca gaggagggca aggaagtatg ggattatgtg 120
acggtccgca aggatgccta catgttcttg tggctctatt atgccacca ctctgcaag 180
aacttctcag aactgccctt ggtcatgttg cttcagggcg gtccaggcgg ttctagcact 240
ggatttgga aacttgagga aattgggcc cttgacagt atctcaaac acggaaaacc 300
acctggctcc aggtgccag tctcctatt gtggataat ccgtgggcac tgggttcagt 360
tatgtgaatg gtagtgggtc ctatgccaag gacctggcta tgggtggctt agacatgatg 420
gttctcctga agaccttctt cagttgccac aaagaattcc agacagttcc attctacatt 480
ttctcagagt cctatgg 497

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<210> 191
<211> 175
<212> DNA
<213> Homo sapiens

<400> 191
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ctacttgagt acaaggattt gagcctgtta cattcactgc tgaatttttag gtccttggaa 120
gatacccagc attcaataga gaccacacaa taaatatatg tcaaataaaa aaaaa 175

<210> 192
<211> 526
<212> DNA
<213> Homo sapiens

<400> 192
agtaaacatt attatTTTTT ttatatttgc aaaggaaaca tatctaattcc ttcctataga 60
aagaacagta ttgctgtaat tccttttctt ttcttctca ttctctctgc cccttaaaag 120
attgaagaaa gagaaacttg tcaactcata tccacgttat ctacgaaagt acataagaat 180
ctatcactaa gtaatgtatc ctccagaatg tgttggttta ccagtgcac cccatattca 240
tcacaaaatt aaagcaagaa gtccatagta atttatttgc taatagtggg tttttaatgc 300
tcagagtttc tgagggtcaa ttttatcttt tcacttacaa gctctatgat cttaaataat 360
ttacttaatg tattttggtg tattttcttc aaattaatat tgggtgtcaa gactatatct 420
aattcctctg atcactttga gaaacaaact ttattaaat gtaaggcact tttctatgaa 480
ttttaaatat aaaaataaat attgttctga ttattactga aaaaaa 526

<210> 193
<211> 553
<212> DNA
<213> Homo sapiens

<220>
<221> unsure
<222> (290)
<223> n=A,T,C or G
<221> unsure
<222> (300)
<223> n=A,T,C or G
<221> unsure
<222> (411)
<223> n=A,T,C or G
<221> unsure
<222> (441)
<223> n=A,T,C or G

<400> 193
tccattgtgg tggaattcgc tctctggtaa aggcgtgcag gtgttgccg cggcctctga 60
gctgggatga gccgtgctcc cgggtggaagc aaggaggccc agccggagcc atggccagta 120
cagtggtagc agttggactg accattgctg ctgcaggatt tgcaggccgt tacgttttgc 180
aagccatgaa gcatatggag cctcaagtaa aacaagtttt tcaaagccta ccaaaatctg 240
ccttcagtgg tggctattat agagggtgggt ttgaacccaa aatgacaaan cggaagcan 300
cattaatact aggtgtaagc cctactgcca ataaaggga aataagagat gctcatcgac 360
gaattatgct tttaaatcat cctgacaaag gaggatctcc ttatatagca nccaaaatca 420
atgaagctaa agatttacta naagggtcaag ctaaaaaatg aagtaaatgt atgatgaatt 480

ttaagttcgt attagtttat gtatatgagt actaagtttt tataataaaa tgcctcagag 540
ctacaatttt aaa 553

<210> 194

<211> 320

<212> DNA

<213> Homo sapiens

<400> 194

cccttcccaa tccatcagta aagaccccat ctgccttgtc catgccgttt cccaacaggg 60
atgtcacttg atatgagaat ctcaaattct aatgccttat aagcattcct tcctgtgtcc 120
attaagactc tgataattgt ctcccccca taggaatttc tcccaggaaa gaaatatatc 180
cccatctccg ttccatatca gaactaccgt ccccgatatt cccttcagag agattaaaga 240
ccagaaaaaa gtgagcctct tcatctgcac ctgtaatagt ttcagttcct attttcttcc 300
attgacccat atttatacct 320

<210> 195

<211> 320

<212> DNA

<213> Homo sapiens

<220>

<221> unsure

<222> (203)

<223> n=A,T,C or G

<221> unsure

<222> (218)

<223> n=A,T,C or G

<400> 195

aagcatgacc tggggaaatg gtcagacctt gtatttgttt tttggccttg aaagtagcaa 60
gtgaccagaa tctgccatgg caacaggctt taaaaaagac ccttaaaaag acactgtctc 120
aactgtgggtg ttagcaccag ccagctctct gtacatttgc tagctttagt ttttctaaga 180
ctgagtaaac ttcttatttt tanaaaagggg aggctggntt gtaactttcc ttgtacttaa 240
ttgggtaaaa gtcttttcca caaaccacca tctattttgt gaactttgtt agtcattctt 300
tatttggtaa attatgaact 320

<210> 196

<211> 357

<212> DNA

<213> Homo sapiens

<220>

<221> unsure

<222> (36)

<223> n=A,T,C or G

<400> 196

atataaaata atacgaaact ttaaaaagca ttggantgtc agtatgttga atcagtagtt 60
tcactttaac tgtaaacaat ttcttaggac accatttggg ctagtcttct tgtaagtgtg 120
aatactacaa aaacttattt atactgttct tatgtcattt gttatattca tagatttata 180
tgatgatatg acatctggct aaaaagaaat tattgcaaaa ctaaccacta tgtacttttt 240
tataaatact gtatggacaa aaaatggcat tttttatatt aaattgttta gctctggcaa 300
aaaaaaaaa ttttaagagc tgggtactaat aaaggattat tatgactgtt aaaaaaa 357

<210> 197
 <211> 565
 <212> DNA
 <213> Homo sapiens

<220>
 <221> unsure
 <222> (27)
 <223> n=A,T,C or G

<400> 197
 tcagctgagt accatcagga tatttanccc ttttaagtgt gttttgggag tagaaaacta 60
 aagcaacaat acttcctctt gacagctttg attggaatgg ggttattaga tcattcacct 120
 tggctctaca ctttttagga tgcttggtga acataacacc acttataatg aacatccctg 180
 gttcctatat tttgggctat gtgggtagga attgttactt gttactgcag cagcagccct 240
 agaaaagtaag cccagggctt cagatctaag ttagtccaaa agctaaatga tttaaagtca 300
 agttgtaatg ctaggcataa gcactctata atacattaaa ttataggccg agcaattagg 360
 gaatgtttct gaaacattaa acttgtattt atgtcactaa aattctaaca caaacttaaa 420
 aaatgtgtct catacatatg ctgtactagg cttcatcatg catttctaaa tttgtgtatg 480
 atttgaatat atgaaagaat ttatacaaga gtgttattta aaattattaa aaataaatgt 540
 atataatttg tacctattgt aaaaa 565

<210> 198
 <211> 484
 <212> DNA
 <213> Homo sapiens

<400> 198
 tatgtaagta ttggtgtctg ctttaaaaaa ggagaccag acttcacctg tcctttttta 60
 acatttgaga acagtgttac tctgagcagt tgggccacct tcaccttacc cgacagctga 120
 ctggttgatg tgtccattgt cgccagtttg gctgttgccc ggacaggaca ggacctccat 180
 tgggcgcagc agcaggtggc aggggtgtgg cttgaggtgg gtggcagcgt ctggctcctc 240
 tctctggtgc tttctgagag ggtctctaaa gcagagtgtg gttggcctgg ggggaaggcag 300
 agcacgtatt tctccctct agtacctctg catttgtgag tgttccctct ggctttctga 360
 agggcagcag actcttgagt atactgcaga ggacatgctt tatcagtagg tcctgagggc 420
 tccaggggct caactgacca agtaacacag aagttggggg atgtggccta tttgggtcgg 480
 aaac 484

<210> 199
 <211> 429
 <212> DNA
 <213> Homo sapiens

<220>
 <221> unsure
 <222> (77)
 <223> n=A,T,C or G
 <221> unsure
 <222> (88)
 <223> n=A,T,C or G
 <221> unsure
 <222> (134)
 <223> n=A,T,C or G
 <221> unsure
 <222> (151)

<223> n=A,T,C or G
 <221> unsure
 <222> (189)
 <223> n=A,T,C or G
 <221> unsure
 <222> (227)
 <223> n=A,T,C or G
 <221> unsure
 <222> (274)
 <223> n=A,T,C or G
 <221> unsure
 <222> (319)
 <223> n=A,T,C or G

<400> 199
 gcttatgttt tttgttttaa cttttgtttt ttaacattta gaatattaca ttttgtatta 60
 tacagtacct ttctcanaca ttttgtanaa ttcatttcgg cagctcacta ggattttgct 120
 gaacattaaa aagngtgata gcgatattag ngccaatcaa atggaaaaaa ggtagtctta 180
 ataaacaana cacaacgttt ttatacaaca tactttaaaa tattaanaaa actccttaat 240
 attgtttcct attaagtatt attctttggg caanattttc tgatgtttt gatttttctt 300
 caatttagca tttgctttng gtttttttct ctatttagca ttctgttaag gcacaaaaac 360
 tatgtactgt atgggaaatg ttgtaaatat taccttttcc acattttaaa cagacaactt 420
 tgaatccaa 429

<210> 200
 <211> 279
 <212> DNA
 <213> Homo sapiens

<400> 200
 gcttttttga ggaattacag ggaagctcct ggaattgtac atggatatct ttatccctag 60
 ggggaaatca aggagctggg caccctaata tctttatgga agtgtttaaa actattttta 120
 ttttattaca agtattacta gtagtggt tctactctaa gatttcaaaa gtgcatttaa 180
 aatcatatcat gttccgcct gcaaatatat tggtattttg gtggagaaaa aaatagtata 240
 ttctacataa aaaattaaag atattaacta agaaaaaa 279

<210> 201
 <211> 569
 <212> DNA
 <213> Homo sapiens

<400> 201
 taggtcagta tttttagaaa ctcttaatag ctcatactct tgataccaaa agcagccctg 60
 attgttaaag cacacacctg cacaagaagc agtgatggtt gcattttacat ttctgggtg 120
 cacaaaaaaa aattctcaaa aagcaaggac ttacgctttt tgcaaaagcct ttgagaagtt 180
 actggatcat aggaagctta taacaagaat ggaagattct taaataactc actttctttg 240
 gtatccagta acagtagatg ttcaaaatat gtagctgatt aataccagca ttgtgaacgc 300
 tgtacaacct tgtggttatt actaagcaag ttactactag cttctgaaaa gtacttcat 360
 aattaatgtt atttatacac tgccttccat gacttttact ttgccctaag ctaatctcca 420
 aaatctgaaa tgctactcca atatcagaaa aaaaggggga ggtggaatta tatttctctg 480
 gattttaaga gtacagagaa tcatgcacat ctctgattag ttcatatatg tctagtgtgt 540
 aataaaagtc aaagatgaac tctcaaaaa 569

<210> 202
 <211> 501

<212> DNA

<213> Homo sapiens

<400> 202

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attaataggc ttaataattg ttggcaagga tccttttgct ttctttggca tgcaagctcc 60
tagcatctgg cagtggggcc aagaaaataa ggtttatgca tgtatgatgg ttttcttctt 120
gagcaacatg attgagaacc agtgtatgtc aacagggtgca tttagagataa ctttaaataga 180
tgtacctgtg tggcttaagc tggaatctgg tcaccttcca tccatgcaac aacttggtca 240
aattcttgac aatgaaatga agctcaatgt gcatatggat tcaatccac accatcgatc 300
atagcaccac ctatcagcac tgaaaactct tttgcattaa gggatcattg caagagcagc 360
gtgactgaca ttatgaaggc ctgtactgaa gacagcaagc tgtagtaga gaccagatgc 420
tttcttggca ggctcgttgt acctcttgga aaacctcaat gcaagatagt gtttcagtc 480
tggcatattt tgaattctg c 501
```

<210> 203

<211> 261

<212> DNA

<213> Homo sapiens

<220>

<221> unsure

<222> (36)

<223> n=A,T,C or G

<221> unsure

<222> (96)

<223> n=A,T,C or G

<400> 203

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gataaaaatga atgagttctg tcatgattca ctattntata acttgcata cctttactgt 120
gttagctctt tgaatgttct tgaaatttta gactttcttt gtaaacaat gatatgtcct 180
tatcattgta taaaagctgt tatgtgcaac agtgtggaga ttccttgtct gatttaataa 240
aatacttaaa cactgaaaaa a 261
```

<210> 204

<211> 421

<212> DNA

<213> Homo sapiens

<400> 204

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agcatctttt ctacaacgtt aaaattgcag aagtagctta tcattaataa acaacaacaa 60
caacaataac aataaatcct aagtgtaaat cagttattct acccctacc aaggatatca 120
gcctgttttt tccctttttt ctctgggaa taattgtggg cttcttccca aatttctaca 180
gcctctttcc tcttctcatg cttgagcttc cctgtttgca cgcattgctg tgcaggactg 240
gcttgtgtgc ttggactcgg ctccagggtg aagcatgctt tcccttgta ctgttggaga 300
aactcaaacc ttcaagccct aggtgtagcc attttgtcaa gtcataact gtatttttgt 360
actggcatta acaaaaaaag aagataaat attgtacat taaacttta taaaacttta 420
a 421
```

<210> 205

<211> 460

<212> DNA

<213> Homo sapiens

<400> 205


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tactctcaca atgaaggacc tggaaatgaaa aatctgtgtc taaacaagtc ctcttttagat 60
tttagtgcaa atccagagcc agcgtcggtt gcctcgagta attctttcat gggtagcttt 120
ggaaaagctc tcaggagacc tcacctagat gcctattcaa gctttggaca gccatcagat 180
tgtcagccaa gagcctttta tttgaaagct cattcttccc cagacttggg ctctgggtca 240
gaggaagatg ggaaagaaag gacagatttt caggaagaaa atcacatttg tacctttaaa 300
cagacttttag aaaactacag gactccaaat tttcagtctt atgacttggg cacatagact 360
gaatgagacc aaaggaaaag cttaacatac tacctcaagg tgaactttta tttaaaagag 420
agagaatctt atgtttttta aatggagtta tgaattttaa 460

```

<210> 206

<211> 481

<212> DNA

<213> Homo sapiens

<400> 206

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tgtggtggaa ttcgggacgc cccagaccc tgactttttc ctgctgggc cgtctctctc 60
tgcggaagca gtgacctctg acccctggtg accttcgctt tgagtgcctt ttgaacgctg 120
gtcccgcggtg acttggtttt ctcaagctct gtctgtccaa agacgctccg gtcgaggtcc 180
cgctgccctt ggggtggatac ttgaacccca gacgcccctc tgtctgtctg tgcctggagg 240
cgcccttccc atctgctctg ccaccggag ctctttccgc cggcgagggt tcccaagccc 300
acctcccgcc ctgctctctg cgggtgtgct ctgggcaagt cctgcacaca caatgcaagt 360
cctggcctcc gcgccgccc gccacgcga gccgtacctg ccgccaactc tgttatttat 420
gggtgaccc cctggaggtg cctcggccc accggggcta tttattgttt aatttatttg 480
t 481

```

<210> 207

<211> 605

<212> DNA

<213> Homo sapiens

<400> 207

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accctttttg gattcagggc tcctcacaat taaaatgagt gtaatgaaac aagggtgaaa 60
tatagaagca tccctttgta tactgttttg ctacttacag tgtacttggc attgctttat 120
ctcattggtt tctcacggtg ggatttctga gatcttaate taagctccaa agttgtctac 180
ttttttgatc ctagggtgct cttttgttt tacagagcag ggtcacttga tttgctagct 240
gggtggcagaa ttggcaccat taccaggtc tgactgacca ccagtcagag gcactttatt 300
tgtatcatga aatgatttga aatcatttga aagcagcgaa gtctgataat gaatgccagc 360
tttcttctgt ctttgataac aaagactcca aatattcttg agaacctgga taaaagtttg 420
aagggtctga ttgggatttg aagacaaaat tgtaggaaat cttacatttt tgcaataaca 480
aacattaatg aaagcaaaac attataaaaag taattttaat tcaccacata cttatcaatt 540
tcttgatgct tccaaatgac atctaccaga tatggttttg tggacatctt tttctgttta 600
cataa 605

```

<210> 208

<211> 655

<212> DNA

<213> Homo sapiens

<400> 208

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ggcgttggtc tggattcccg tcgtaactta aagggaact ttcacaatgt ccggagccct 60
tgatgtcctg caaatgaagg aggaggatgt ccttaagttc cttgcagcag gaaccactt 120
agggtggcacc aatettgact tccagatgga acagtacatc tataaaagga aaagtgatgg 180
catctatatc ataaatctca agaggacctg ggagaagctt ctgctggcag ctctgtgcaat 240
tgttgccatt gaaaacctg ctgatgtcag tgttatatcc tccaggaata ctggccagag 300
ggctgtgctg aagtttgctg ctgccactgg agccactcca attgctggcc gcttcaactc 360

```

```

tggaaccttc actaaccaga tccaggcagc ctcccgagg ccacggcttc ttgtgggttac 420
tgaccccgagg gctgaccacc agcctctcac ggaggcatct tatgttaacc tacctaccat 480
tgcgctgtgt aacacagatt ctctctgcg ctatgtggac attgccatcc catgcaacaa 540
caaggaggct cactcagtgg gtttgatgtg gtggatgctg gctcggaag ttctgcgcat 600
gcgtggcacc atttcccggtg aacacccatg ggaggatcat cctgatctgt acttc 655

```

<210> 209

<211> 621

<212> DNA

<213> Homo sapiens

<400> 209

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catttagaac atggttatca tccaagacta ctctaccctg caacattgaa ctccaagag 60
caaatccaca ttctcttga gttctgcagc ttctgtgtaa atagggcagc tgcgtctat 120
gccgtagaat cacatgatct gaggaccatt catggaagct gctaaatagc ctagtctggg 180
gagtccttcca taaagttttg catggagcaa acaaacagga ttaaactagg ttgtgttct 240
tcagccctct aaaagcatag ggcttagcct gcaggcttcc ttgggcttcc tctgtgtgtg 300
tagttttgta aacactatag catctgttaa gatccagtgt ccatggaaac cttcccat 360
gccgtgactc tggactatat cagtttttgg aaagcagggt tcctctgcct gctaacaagc 420
ccacgtggac cagtctgaat gtctttcctt tacacctatg tttttaata gtcaaacctc 480
aagaaacaat ctaaacaagt ttctgttgca tatgtgtttg tgaacttgta tttgtattta 540
gtaggcttct atattgcatt taacttgttt ttgtaactcc tgattcttcc ttttcggata 600
ctattgatga ataaagaat t 621

```

<210> 210

<211> 533

<212> DNA

<213> Homo sapiens

<220>

<221> unsure

<222> (20)

<223> n=A,T,C or G

<221> unsure

<222> (21)

<223> n=A,T,C or G

<221> unsure

<222> (61)

<223> n=A,T,C or G

<400> 210

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cgcttgggg agccggcggn ngagtccggg acgtggagac ccgggggtccc ggcagccggg 60
nggcccgcgg gccaggggtg gggatgcacc gccgcggggt gggagctggc gccatcgcca 120
agaagaaact tgagaggcc aagtataagg agcaggggac ggtcttggtc gaggaccagc 180
tagcccagat gtcaaagcag ttggacatgt tcaagaccac cctggaggaa ttgccagca 240
aacacaagca ggagatccgg aagaatcctg agttccgtgt gcagttccag gacatgtgtg 300
caaccattgg cgtggatccg ctggcctctg gaaaaggatt ttggtctgag atgctgggcg 360
tgggggactt ctattacgaa ctaggtgtcc aaattatcga agtgtgcctg gcgtgaagc 420
atcggaatgg aggtctgata actttggagg aactacatca acaggtgttg aagggaaggg 480
gcaagttcgc ccaggatgtc agtcaagatg acctgatcag agccatcaag aaa 533

```

<210> 211

<211> 451

<212> DNA

<213> Homo sapiens

<400> 211
ttagcttgag ccgagaacga ggcgagaaag ctggagaccg aggagaccgc ctagagcgga 60
gtgaacgggg aggggaccgt ggggaccggc ttgatcgtgc gcggacacct gctaccaagc 120
ggagcttcag caaggaagtg gaggagcgga gtagagaacg gccctcccag cctgaggggc 180
tgcgcaaggc agctagcctc acggaggatc gggaccgtgg gcgggatgcc gtgaagcgag 240
aagctgccct acccccagtg agccccctga agcggtctct ctctgaggag gagttagaga 300
agaaatccaa ggctatcatt gaggaatatt tccatctcaa tgacatgaaa gaggcagtcc 360
agtgcgtgca ggagctggcc tcacctctct tgctcttcat ctttgtacgg catggtgtcg 420
agtctacgct ggagcgagcgt gccattgctc g 451

<210> 212
<211> 471
<212> DNA
<213> Homo sapiens

<220>
<221> unsure
<222> (54)
<223> n=A,T,C or G

<400> 212
gtgattattc ttgatcaggg agaagatcat ttagatttgt tttgcattcc ttanaatgga 60
gggcaacatt ccacagctgc cctggctgtg atgagtgtcc ttgcaggggc cggagtagga 120
gcaactgggt gggggcgga ttgggggttac tcgatgtaag ggattccttg ttgttgtgtt 180
gagatccagt gcagttgtga tttctgtgga tccagcttg gttccaggaa ttttgtgtga 240
ttggcttaaa tccagttttc aatcttcgac agctgggctg gaacgtgaac tcagtagctg 300
aacctgtctg acccggtcac gttcttgat cctcagaact ctttgcctct gtcgggtgg 360
gggtgggaac tcacgtgggg agcggtggct gagaaaatgt aaggattctg gaatacatat 420
tccatgggac ttctcttccc tctctgctt cctcttttcc tgctccctaa c 471

<210> 213
<211> 511
<212> DNA
<213> Homo sapiens

<220>
<221> unsure
<222> (27)
<223> n=A,T,C or G
<221> unsure
<222> (63)
<223> n=A,T,C or G
<221> unsure
<222> (337)
<223> n=A,T,C or G
<221> unsure
<222> (442)
<223> n=A,T,C or G

<400> 213
ctaattagaa acttgctgta ctttttnttt tcttttaggg gtcaaggacc ctctttatag 60
ctnccatttg cctacaataa attattgcag cagtttgcaa tactaaaata ttttttatag 120
actttatatt ttctcttttg ataaagggat gctgcatagt agagttggtg taattaaact 180
atctcagcgg ttctcctgct ttcccttctg ctccatatgc ctcattgtcc ttccagggag 240

```

ctcttttaat cttaaagttc tacatttcat gctcttagtc aaattctgtt accttttttaa 300
taactcttcc cactgcatat ttccatcttg aattggnggt tctaaattct gaaactgtag 360
ttgagataca gctattttaat atttctggga gatgtgcac cctcttcttt gtggttgccc 420
aaggttgttt tgcgtaactg anactccttg atatgcttca gagaatttag gcaaactg 480
ggcatggccg tgggagtact gggagtaaaa t 511

```

<210> 214

<211> 521

<212> DNA

<213> Homo sapiens

<400> 214

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agcattgcc aataatccct aattttccac taaaaatata atgaaatgat gttaagcttt 60
ttgaaaagtt taggttaaac ctactgttgt tagattaatg tatttggtgc ttccctttat 120
ctggaatgtg gcattagctt ttttatttta accctcttta attcttattc aattccatga 180
cttaagggtg gagagctaaa cactgggatt ttgggataac agactgacag ttttgcataa 240
ttataatcgg cattgtacat agaaaggata tggctacctt ttgttaaate tgcactttct 300
aaatatcaaa aaagggaat gaagtataaa tcaatttttg tataatctgt ttgaaacatg 360
agtttttatt gcttaatttt agggctttgc cccttttctg taagtctctt gggatcctgt 420
gtagaagctg ttctcattaa acaccaaaca gttaagtcca ttctctggtg ctagtacaa 480
attcgggttc atattctact taacaattta aataaactga a 521

```

<210> 215

<211> 381

<212> DNA

<213> Homo sapiens

<220>

<221> unsure

<222> (17)

<223> n=A,T,C or G

<221> unsure

<222> (20)

<223> n=A,T,C or G

<221> unsure

<222> (60)

<223> n=A,T,C or G

<221> unsure

<222> (61)

<223> n=A,T,C or G

<221> unsure

<222> (365)

<223> n=A,T,C or G

<400> 215

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gagcggagag cggaccngtn agagccctga gcagcccccac cgccgcccgc ggcctagttn 60
ncatcacacc ccgggaggag ccgcagctgc cgcagccggc cccagtcacc atcacgcaa 120
ccatgagcag cgaggccgag acccagcagc cgcccgcgc ccccccgcc gccccgccc 180
tcagcgccgc cgacaccaag cccggcacta cgggcagcgg cgcagggagc ggtggcccgg 240
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<210> 216

<211> 425

<212> DNA

<213> Homo sapiens

<400> 216

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aacaggccaa tctgaagggt actcctgtgt tgctgcagaa tgtcagatat tttggatgtt 180
gcataagagt cctatttgcc ccagttaatt caacttttgt ctgcctgttt tgtggactgg 240
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aattgacaat atatatgcat gtgtttaaac caaatccaga aagcttaaac aatagagctg 360
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<210> 217

<211> 181

<212> DNA

<213> Homo sapiens

<400> 217

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actttggctt tttcagtgga agaatatgtt gaaggtttca ttttgttcta gaaaaaaaaa 180
a                                             181

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<210> 218

<211> 405

<212> DNA

<213> Homo sapiens

<400> 218

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gcgctgggct gtttttagtgc caggctgcgg tgggcagcca tgagaacaaa acctcttctg 180
tatttttttt ttccattagt aaaacacaag acttcagatt cagccgaatt gtggtgtctt 240
acaaggcagg cctttcttac aggggggtgga gagaccagcc tttcttcctt tggtaggaat 300
ggcctgagtt ggcgttgttg gcaggctact ggtttgatg atgtattagt agagcaaccc 360
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<211> 216

<212> DNA

<213> Homo sapiens

<220>

<221> unsure

<222> (207)

<223> n=A,T,C or G

<221> unsure

<222> (210)

<223> n=A,T,C or G

<400> 219

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tcaattgtaa acttcttgtt aagactgtta cgtttctatt gcttttgtat gggatattgc 180

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aaaaataaaa aggaagaac cctcttnaan aaaaaa

216

<210> 220

<211> 380

<212> DNA

<213> Homo sapiens

<400> 220

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gcacccaag gactcagaag atgattttaa cagttcagaa cagatgtgtg caatattggt 240
gcatgtaata atgttgagtg gcagtcaaaa gtcattgatt ttatcttagt tcttcattac 300
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gtaagtcttt gacaaaaaaa 380

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<211> 398

<212> DNA

<213> Homo sapiens

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cccagccccg tttcctttta ttttgagct aatgccagct gcgtgtctag ttttgagtgc 240
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gtttctgtgg gagcagtgt caccactct tctgtatat tgccttttg ctggaaaatg 360
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<211> 301

<212> DNA

<213> Homo sapiens

<220>

<221> unsure

<222> (49)

<223> n=A,T,C or G

<221> unsure

<222> (64)

<223> n=A,T,C or G

<400> 222

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gatgacttta ggatttgcat ttttcctttt attgcctcat ttcttgtgac gccttggttg 240
ggagggaaat ctgtttattt tttcctacaa ataaaaagct aagattctat atcgcaaaaa 300
a 301

<210> 223

<211> 200

<212> DNA

<213> Homo sapiens

<400> 223

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 agatttctac aggagacagt ggttttattt ggattgtctt ctgtaatagg tttcaataaa 180
 gctggatgaa cttaaaaaaa 200

<210> 224

<211> 385

<212> DNA

<213> Homo sapiens

<400> 224

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 tctccaacac cagcaagccc taaccagggc cctcctccac aagttccagt atctcctgga 180
 ccaccaaagg acagttctgc ccttggtgga cccccagaaa ggactgttac tccagcccta 240
 tcatcaaagt tgttaccaag acatcttgga tccccgcta cttcagtgcc tggaatgggt 300
 aaacagagca cttaatgtta tttacagttt atattgtttt ctctggttac caataaaacg 360
 ggccattttc aggtggtaaa aaaaa 385

<210> 225

<211> 560

<212> PRT

<213> Homo sapien

<400> 225

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Leu	Pro	Leu	Asp	Ala	Ala	Lys	Arg	Phe	His	Asp	Val	Leu	Gly	Asn	Glu	20	25	30	
Arg	Pro	Ser	Ala	Tyr	Met	Arg	Glu	His	Asn	Gln	Leu	Asn	Gly	Trp	Ser	35	40	45	
Ser	Asp	Glu	Asn	Asp	Trp	Asn	Glu	Lys	Leu	Tyr	Pro	Val	Trp	Lys	Arg	50	55	60	
Gly	Asp	Met	Arg	Trp	Lys	Asn	Ser	Trp	Lys	Gly	Gly	Arg	Val	Gln	Ala	65	70	75	80
Val	Leu	Thr	Ser	Asp	Ser	Pro	Ala	Leu	Val	Gly	Ser	Asn	Ile	Thr	Phe	85	90	95	
Ala	Val	Asn	Leu	Ile	Phe	Pro	Arg	Cys	Gln	Lys	Glu	Asp	Ala	Asn	Gly	100	105	110	
Asn	Ile	Val	Tyr	Glu	Lys	Asn	Cys	Arg	Asn	Glu	Ala	Gly	Leu	Ser	Ala	115	120	125	
Asp	Pro	Tyr	Val	Tyr	Asn	Trp	Thr	Ala	Trp	Ser	Glu	Asp	Ser	Asp	Gly	130	135	140	
Glu	Asn	Gly	Thr	Gly	Gln	Ser	His	His	Asn	Val	Phe	Pro	Asp	Gly	Lys	145	150	155	160
Pro	Phe	Pro	His	His	Pro	Gly	Trp	Arg	Arg	Trp	Asn	Phe	Ile	Tyr	Val	165	170	175	
Phe	His	Thr	Leu	Gly	Gln	Tyr	Phe	Gln	Lys	Leu	Gly	Arg	Cys	Ser	Val	180	185	190	
Arg	Val	Ser	Val	Asn	Thr	Ala	Asn	Val	Thr	Leu	Gly	Pro	Gln	Leu	Met	195	200	205	
Glu	Val	Thr	Val	Tyr	Arg	Arg	His	Gly	Arg	Ala	Tyr	Val	Pro	Ile	Ala				

210	215	220
Gln Val Lys Asp Val Tyr Val Val Thr Asp Gln Ile Pro Val Phe Val		
225	230	235
Thr Met Phe Gln Lys Asn Asp Arg Asn Ser Ser Asp Glu Thr Phe Leu		240
	245	250
Lys Asp Leu Pro Ile Met Phe Asp Val Leu Ile His Asp Pro Ser His		255
	260	265
Phe Leu Asn Tyr Ser Thr Ile Asn Tyr Lys Trp Ser Phe Gly Asp Asn		270
	275	280
Thr Gly Leu Phe Val Ser Thr Asn His Thr Val Asn His Thr Tyr Val		285
	290	295
Leu Asn Gly Thr Phe Ser Leu Asn Leu Thr Val Lys Ala Ala Ala Pro		300
305	310	315
Gly Pro Cys Pro Pro Pro Pro Pro Pro Arg Pro Ser Lys Pro Thr		320
	325	330
Pro Ser Leu Gly Pro Ala Gly Asp Asn Pro Leu Glu Leu Ser Arg Ile		335
	340	345
Pro Asp Glu Asn Cys Gln Ile Asn Arg Tyr Gly His Phe Gln Ala Thr		350
	355	360
Ile Thr Ile Val Glu Gly Ile Leu Glu Val Asn Ile Ile Gln Met Thr		365
	370	375
Asp Val Leu Met Pro Val Pro Trp Pro Glu Ser Ser Leu Ile Asp Phe		380
385	390	395
Val Val Thr Cys Gln Gly Ser Ile Pro Thr Glu Val Cys Thr Ile Ile		400
	405	410
Ser Asp Pro Thr Cys Glu Ile Thr Gln Asn Thr Val Cys Ser Pro Val		415
	420	425
Asp Val Asp Glu Met Cys Leu Leu Thr Val Arg Arg Thr Phe Asn Gly		430
	435	440
Ser Gly Thr Tyr Cys Val Asn Leu Thr Leu Gly Asp Asp Thr Ser Leu		445
	450	455
Ala Leu Thr Ser Thr Leu Ile Ser Val Pro Asp Arg Asp Pro Ala Ser		460
465	470	475
Pro Leu Arg Met Ala Asn Ser Ala Leu Ile Ser Val Gly Cys Leu Ala		480
	485	490
Ile Phe Val Thr Val Ile Ser Leu Leu Val Tyr Lys Lys His Lys Glu		495
	500	505
Tyr Asn Pro Ile Glu Asn Ser Pro Gly Asn Val Val Arg Ser Lys Gly		510
	515	520
Leu Ser Val Phe Leu Asn Arg Ala Lys Ala Val Phe Phe Pro Gly Asn		525
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Gln Glu Lys Asp Pro Leu Leu Lys Asn Gln Glu Phe Lys Gly Val Ser		540
545	550	555
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<212> PRT

<213> Homo sapien

<400> 226

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5

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<211> 9

<212> PRT

<213> Homo sapien

<400> 227

Phe Leu Leu Asn Asp Asn Leu Thr Ala

1

5

<210> 228

<211> 9

<212> PRT

<213> Homo sapien

<400> 228

Leu Leu Gly Asn Cys Leu Pro Thr Val

1

5

<210> 229

<211> 10

<212> PRT

<213> Homo sapien

<400> 229

Lys Leu Leu Gly Asn Cys Leu Pro Thr Val

1

5

10

<210> 230

<211> 10

<212> PRT

<213> Homo sapien

<400> 230

Arg Leu Thr Gly Gly Leu Lys Phe Phe Val

1

5

10

<210> 231

<211> 9

<212> PRT

<213> Homo sapien

<400> 231

Ser Leu Gln Ala Leu Lys Val Thr Val

1

5

<210> 232

<211> 20

<212> PRT

<213> Homo sapiens

<400> 232

Ala Gly Ala Asp Val Ile Lys Asn Asp Gly Ile Tyr Ser Arg Tyr Phe

5

10

15

Phe Ser Phe Ala

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<213> Homo sapiens

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Asn His Ser Pro Ser
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<213> Homo sapiens

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Asp Pro Asp Gly
20

<210> 235
<211> 20
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<213> Homo sapiens

<400> 235
Leu Gln Ser Ala Val Ser Asn Ile Ala Gln Ala Pro Leu Phe Ile Pro
5 10 15

Pro Asn Ser Asp
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<210> 236
<211> 20
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<213> Homo sapiens

<400> 236
Ile Gln Asp Asp Phe Asn Asn Ala Ile Leu Val Asn Thr Ser Lys Arg
5 10 15

Asn Pro Gln Gln
20

<210> 237

<211> 21
<212> PRT
<213> Homo sapiens

<400> 237
Arg Asn Ser Leu Gln Ser Ala Val Ser Asn Ile Ala Gln Ala Pro Leu
5 10 15
Phe Ile Pro Pro Asn
20

<210> 238
<211> 20
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<213> Homo sapiens

<400> 238
Thr His Glu Ser His Arg Ile Tyr Val Ala Ile Arg Ala Met Asp Arg
5 10 15
Asn Ser Leu Gln
20

<210> 239
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<400> 239
Arg Asn Pro Gln Gln Ala Gly Ile Arg Glu Ile Phe Thr Phe Ser Pro
5 10 15
Gln Ile Ser Thr
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<210> 240
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<400> 240
Gly Gln Ala Thr Ser Tyr Glu Ile Arg Met Ser Lys Ser Leu Gln Asn
5 10 15
Ile Gln Asp Asp Phe
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<210> 241
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<213> Homo sapiens

<400> 241

Glu Arg Lys Trp Gly Phe Ser Arg Val Ser Ser Gly Gly Ser Phe Ser
5 10 15Val Leu Gly Val
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<210> 242

<211> 20

<212> PRT

<213> Homo sapiens

<400> 242

Gly Ser His Ala Met Tyr Val Pro Gly Tyr Thr Ala Asn Gly Asn Ile
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<212> PRT

<213> Homo sapiens

<400> 243

Val Asn His Ser Pro Ser Ile Ser Thr Pro Ala His Ser Ile Pro Gly
5 10 15Ser His Ala Met
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<210> 244

<211> 20

<212> PRT

<213> Homo sapiens

<400> 244

Ala Val Pro Pro Ala Thr Val Glu Ala Phe Val Glu Arg Asp Ser Leu
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<211> 20

<212> PRT

<213> Homo sapiens

<400> 245

Lys Pro Gly His Thr Tyr Thr Leu Asn Asn Thr His His Ser Leu

117

5

10

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Gln Ala Leu Lys
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<210> 246
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<400> 246
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Pro Gly His Trp
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<210> 247
<211> 20
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<400> 247
Leu His Phe Pro His Pro Val Met Ile Tyr Ala Asn Val Lys Gln Gly
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Phe Tyr Pro Ile
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<210> 248
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<400> 248
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Gly Ala Asp Val
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<210> 249
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<400> 249
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Glu Thr Gly Asp

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<210> 250
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<400> 250
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Leu Thr Phe Arg
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<210> 251
 <211> 20
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 <213> Homo sapiens

<400> 251
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 5 10 15

Val Pro Pro Ala
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<210> 252
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 <212> PRT
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<400> 252
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 Trp Arg Pro Val Lys Ala Ser Asp Gly Asp Tyr Tyr Thr Leu Ala Val
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 Pro Met Gly Asp Val Pro Met Asp Gly Ile Ser Val Ala Asp Ile Gly
 35 40 45
 Ala Ala Val Ser Ser Ile Phe Asn Ser Pro Glu Glu Phe Leu Gly Lys
 50 55 60
 Ala Val Gly Leu Ser Ala Glu Ala Leu Thr Ile Gln Gln Tyr Ala Asp
 65 70 75 80
 Val Leu Ser Lys Ala Leu Gly Lys Glu Val Arg Asp Ala Lys Ile Thr
 85 90 95
 Pro Glu Ala Phe Glu Lys Leu Gly Phe Pro Ala Ala Lys Glu Ile Ala
 100 105 110
 Asn Met Cys Arg Phe Tyr Glu Met Lys Pro Asp Arg Asp Val Asn Leu
 115 120 125
 Thr His Gln Leu Asn Pro Lys Val Lys Ser Phe Ser Gln Phe Ile Ser
 130 135 140
 Glu Asn Gln Gly Ala Phe Lys Gly Met
 145 150

<210> 253
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 <213> Homo sapien

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<210> 255

<211> 401

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(401)

<223> n = A,T,C or G

<400> 255

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agtccanagg acggagaaga cgaggaagag	gaggagcagt	tggttctggg	ggaattatca	120
ggaattattg attcagactt cctctcaaaa	tgtgaaaata	aatgcaagg	tttgggcatt	180
gacactgaga ggcccattct gcaagtggac	agctgtgtct	ttgctgggga	gtatgaagac	240
actctangga cctgtgttat atttgaagaa	aatgntnaac	atgctgatac	agaaggcaat	300
aataaaacag tgctaaaata taaatgccat	acaatgaaga	agctcagcat	gacaagaact	360
ctcctgacag agaagaagga aggagaagaa	aacatangtg	g		401

<210> 256

<211> 401

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(401)

<223> n = A,T,C or G

<400> 256

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gggcgggggt cgcggccgng gacggggcgc	gggcnangc	cgnganctc	gcggangcaa	120
ggccgaggat aaggagtga tgcccgtcac	caacttgggc	cgcttgncca	aggacatgaa	180
nancaagccc ctgnaggaga tctatntctt	cttccctgcc	ccattaagga	atcaagagat	240
catttgattt cttcctgggg gcctctctca	aggatnagg	ttttgaagat	tatgccagt	300
canaaaannan acccggttgc ccngtccatc	tncacccaac	ncttccaagg	gcnatTTTTg	360
tttaggcctc attncgggg ggaaccttaa	cccaatttgg	g		401

<210> 257

<211> 401

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(401)

<223> n = A,T,C or G

<400> 257

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ctctcagccc tgaggtatac agaatcattt	gcctcagact	gctgttggat	tttaaaattt	120
ttaaaataac tgctaagtaa tttgtatgt	cttctccac	actatcaata	tgctgtctc	180
taacaggctc ccactttct tttaatgtgc	tgttatgagc	tttggacatg	agataaccgt	240
gcctgttcag agtgtctaca gtaagagctg	gacaaactct	ggagggacac	agtctttgag	300
acagctcttt tggttgcttt ccactttct	gaaaggttca	cagtaacctt	ctagataata	360
gaaactccca gttaaagcct angctancaa	ttttttttag	t		401

<210> 258

<211> 401

<212> DNA

<213> Homo sapien

<400> 258

ggagcgctag gtcggtgtac gaccgagatt aggggtgcgtg ccagctccgg gaggccgcgg	60
tgagggggccg ggcccaagct gccgacccga gccgatcgtc agggtcgcca gcgcctcagc	120
tctgtggagg agcagcagta gtcggagggt gcaggatatt agaaatggct actccccagt	180
caattttcat ctttgcaatc tgcattttta tgataacaga attaattctg gcctcaaaaa	240
gctactatga tatcttaggt gtgccaaaat cggcatcaga gcgccaaatc aagaaggcct	300
ttcacaagtt ggccatgaag taccaccctg acaaaaataa gacccagatg ctgaagcaaa	360
attcagagag attgcagaag catatgaaac actctcagat g	401

<210> 259

<211> 401

<212> DNA

<213> Homo sapien

<400> 259

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ctccagaata ttgtgggttt gatcatcaat gcagtcatgt taggctgcat ttcatgaaa	120
acagctcagg ctacagaag ggcagaaact ttgattttca gccgccatgc tgtgattgcc	180
gtccgaaatg gcaagctgtg cttcatgttc cgagtgggtg acctgaggaa aagcatgac	240
attagtgcct ctgtgcgcat ccagggtggtc aagaaaacaa ctacacctga aggggagggtg	300
gttcctatc accaactgga cattcctgtt gataacccaa tcgagagcaa taacattttt	360
ctgttggtccc ctttgatcat ctgccacgtg attgacaagc g	401

<210> 260

<211> 363

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (363)

<223> n = A,T,C or G

<400> 260

aggaganang gagggggana tgaataggga tggagaggga natagtggat gagcagggca	60
caggagagg aancagaaag gagaggcaag acaggagagac acacancaca nangangana	120
cagggtggggg ctgggggtggg gcatggagag ccttttngat cncacaggcc accctgctct	180
cgctggngctg ttgaaaccca ctccatggct tectgccact gcagttggggc ccagggtggtg	240
cttattnctg gaatgcaagt ggctgtggct tggagcctcc cctctggnnn anggaaannn	300
attgctccct tatctgcttg gaatatctga gtttttccan cccggaaata aaacacacac	360
aca	363

<210> 261

<211> 401

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (401)

<223> n = A,T,C or G

<400> 261

cggtctccg ccgtctccc ggggtttcgg ggcacttggg tcccacagtc tggctcgtct	60
tcacctccc ctgacctgag tagtcgccat ggcacagggt ctcagaggca ctgngactga	120

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cttccctgga tttgatgagc gggctgatgc anaaactctt cggaaggcta tgaaaggctt 180
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ggaaatctct gcagctttta agactctgtt tggcagggat cttctggatg acctgaaatc 300
agaactaact ggaaaatttg aaaaattaat tgtggctctg atgaaaccct ctcggcttta 360
tgatgcttat gaactgaaac atgccttgaa gggagctgga a 401

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<210> 262
<211> 401
<212> DNA
<213> Homo sapien

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<220>
<221> misc_feature
<222> (1)...(401)
<223> n = A,T,C or G

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<400> 262
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agtttataac atgaagaata ttgtaccatt atacattttc attctcgatc tcataagaaa 180
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tcaactcaaa aattatgntg catagtttta ttttgaattt aggttttggg actacttttt 300
tccancttca atgagaaaat aaaatctaca actcaggagt tactacagaa gttctaanta 360
tttttttgct aannagcnaa aaatataaac atatgaaaat g 401

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<210> 263
<211> 401
<212> DNA
<213> Homo sapien

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<220>
<221> misc_feature
<222> (1)...(401)
<223> n = A,T,C or G

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<400> 263
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cgggcggttg cggttagggc ggcgcggaat aaaggggccc cgcccggttg atgcggtgac 180
cactgcggca ggcccaggag ctgagtgggc cccggccctc agcccgtecc gncggaccgg 240
ctttcctcaa ctctccatct tctctgccc accgagatcg ccgaggcggn ctcaggctcc 300
ctanccctt ccccgtecc tcccncccc ogccccgcc ccggggggccg ccgccaccgg 360
cctccacca tggctctgaa ganaatccac aaggaattga a 401

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<210> 264
<211> 401
<212> DNA
<213> Homo sapien

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<400> 264
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aagccacccc ctggcagaaa cttcagctgt gtgttctgga atactcacgt gagggaaactt 120
actttggcca gcattgacct tcaaagtcag atggaacca ggacctatcc aacttggctg 180
cttcacattt tcatccctc ctgcatcatt gctttcattt tcatagccac agtgatagcc 240
ctaagaaaac aactctgtca aaagctgtat tcttcaaaag acacaacaaa aagacctgtc 300

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accacaacaa agagggaagt gaacagtgt gtgaatctga acctgtgggc ttgggagcca 360
gggtgacctg atatgacatc taaagaagct tctggactct g 401

<210> 265

<211> 271

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(271)

<223> n = A,T,C or G

<400> 265

gccacttcct gtggacatgg gcagagcgct gctgccagtt cctggtagcc ttgaccacna 60
cgctgggggg tctttgtgat ggtcatgggt ctcatttgca cttgggggtg tgggattcaa 120
gttagaagtt tctagatctg gccgggcgca gtggctcaca cctgtaatcc cagcacttta 180
ggaggctgag gcaggcgat catgaggtca ggagatcgag accgtcctgg ctaacacagt 240
gaaaccccgct ctctactaaa aatacaaaaa a 271

<210> 266

<211> 401

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(401)

<223> n = A,T,C or G

<400> 266

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gacagcaaaa ttttcatgaa atgtaaaata tttttatagt ttgttcatac tatatgaggt 120
tctattttta atgactttct ggatttttaa aaatttcttt aaatacaatc atttttgtaa 180
tattttatgt atgcttatga tctagataat tgcagaatat cattttatct gactctgtct 240
tcataagaga gctgtggcgg aattttgaac atctgttata gggagtgatc aaattagaag 300
gcaatgtgga aaaacaatc tgggaagat ttctttatat gaagtcctcg ccactagcca 360
gccatcctaa ttgatgaaag ttatctgttc acaggcctgc a 401

<210> 267

<211> 401

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(401)

<223> n = A,T,C or G

<400> 267

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catgcagctg ttcccgcgag gctgtttga ggacgcgctg ccgcccacg tgctgaggag 180
ccaggtgtac agccttgtgc ctgacaggac cgtggccgac cggcagctga aggagcttca 240
agagcanggg gagacaaaat cgtccagctg ggcttcnact tggatgccca tggaanttat 300

tctttcnctt ganggactta cnnngggaccc aagaancctt tncaaggggc ccttngtgga 360
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<210> 268

<211> 223

<212> DNA

<213> Homo sapien

<400> 268

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 gaatcaagta gtcacgcact ttttctgttc atttttctaa aaagtaaata tacaaatgtt 180
 ttgttttttg ttttttttgt ttgtttgttt ctgttttttt ttt 223

<210> 269

<211> 401

<212> DNA

<213> Homo sapien

<400> 269

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 gtttattttt atttaaagt caatagttgt tttttaaagt ccaaatcaga ggtgcaggcc 180
 accagttaaa tgccgtctat caggttttgt gccttaagag actacagagt caaagctcat 240
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 cattttgtct cattgttttc tttgacataa ctaggatcca t 401

<210> 270

<211> 401

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (401)

<223> n = A, T, C or G

<400> 270

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 tgtttgagcc ccatggcact gagctggaat ctgagggtct tgttccaagg atgtgatgat 180
 gtggggagaat gttctttgaa agagcagaaa tccagtctgc atggaaacag cctgtagagn 240
 agaagtttcc agtgataagt gttcactgtt ctaaggaggt acaccacagc tacctgaatt 300
 ttcccaaaat gagtgttct gtgcgttaca actggccttt gtacttgact gtgatgactt 360
 tgttttttct tttcaattct anatgaacat gggaaaaaat g 401

<210> 271

<211> 329

<212> DNA

<213> Homo sapien

<400> 271

ccacagcctc caagtcagg ggggtggagt cccagagctg cacagggttt ggcccaagtt 60
 tctaaggag gcaacttctc ccctcgccca tcagtgccag ccctgtctgg ctggtgctg 120

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agccccctcag acagccccct gccccgcagg cctgccttct cagggacttc tgcggggcct      180
gaggcaagcc atggagtgag acccaggagc cggacacttc tcaggaaatg gcttttccca      240
acccccagcc cccacccggt ggttcttctc gttctgtgac tgtgtatagt gccaccacag      300
cttatggcat ctcattgagg acaaaaaaa      329

```

<210> 272

<211> 401

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(401)

<223> n = A,T,C or G

<400> 272

```

nggctgntaa cntcggaggt nacttctctg actatcctgg agacccctc cgcttccacg      60
nncatnatat cntcatngc tgggccentn angacacnat cccactccaa cacctgngng      120
atgctggncc cctnggaacc ancntcagaa ngaccctgnt cntntgtntt cgcgaanctg      180
aagmnaangc gggntacacc tncntgcant ggnccacnct gcnggggaact ntacacacct      240
acgggatgtg gctgcgccan gagccaagag cntttctgga tgattcccca gcctcttgnn      300
agggantcta caacattgct nnttaccttt ntccnncngc nntnnttga ntacaggngn      360
tnntaacact acatcttttt tactgcncn tncctgtgtg g      401

```

<210> 273

<211> 401

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(401)

<223> n = A,T,C or G

<400> 273

```

cagcaccatg aagatcaaga tcatcgcacc cccagagcgc aagtactcgg tgtggatcgg      60
tggtccatc ctggcctcac tgtccacctt ccagcagatg tggattagca agcaggagta      120
cgacgagtcg gggccctcca tcgtccaccg caaatgcttc taaacggact cagcagatgc      180
gtagcatttg ctgcatgggt taattgagaa tagaaatttg cccctggcaa atgcacacac      240
ctcatgctag cctcacgaaa ctggaataag ccttcgaaaa gaaattgtcc ttgaagcttg      300
tatctgatat cagcactgga ttgtagaact tgttgctgat tttgaccttg tattgaagtt      360
aactgttccc cttggtatta acgtgtcagg gctgagtnt c      401

```

<210> 274

<211> 401

<212> DNA

<213> Homo sapien

<400> 274

```

ccaccacac ccaccgcgc ctcgttcgcc tcttctcgg gagccagtc gcgccaccgc      60
cgccgcccag gccatcgcca cctcgcgag ccatgtccac caggtcggtg tctcgtcct      120
cctaccgcag gatgttcggc ggccggggca ccgcgagccg gccgagctcc agcgggagct      180
acgtgactac gtccaccgc acctacagcc tgggcagcgc gctgcgcccc agcaccagcc      240
gcagcctcta cgctcgtcc ccggggcgcg tgtatgccac gcgctcctct gccgtgcgcc      300
tgccgagcag cgtgcccggg gtgcggctcc tgcaggactc ggtggacttc tcgctggccg      360

```

acgccatcaa caccgagttc aagaacaccc gcaccaacga g 401

<210> 275
 <211> 401
 <212> DNA
 <213> Homo sapien

<400> 275
 ccacttccac cactttgttg agcagtgcct tcagcgcaac ccggatgcca ggtatccctg 60
 ctggcctggg cctgggcttc gggagagcag aggggtgctca ggagggttaag gccaggggtg 120
 gaagggaactt acctcccaaa ggttctgcag gggaatctgg agctacacac aggagggatc 180
 agctcctggg tgtgtcagag gccagcctgg ggagctctgg ccactgcttc ccatgagctg 240
 agggagaggg agaggggacc cgaggctgag gcataagtgg caggatttcg ggaagctggg 300
 gacacggcag tgatgctgag gtctctcttc ccctttccct ccaggcccag tgccagcacc 360
 ctctgaacc actctttctt caagcagatc aagcgacgtg c 401

<210> 276
 <211> 401
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(401)
 <223> n = A,T,C or G

<400> 276
 tctgatattg ntacccttga gccacctaag ttagaagaaa ttggaaatca agaagttgtc 60
 attgttgaag aagcacagag ttcagaagac tttaacatgg gctcttcctc tagcagccag 120
 tatactttct gtcagccaga aactgtattt tcctctcagc ctagtgtatga tgaatcaagt 180
 agtgatgaaa ccagtaatca gccagtcctt gccttttagac gacgccgtgc taggaagaag 240
 accgtttctg cttcagaatc tgaagaccgg ctagtgtggtg aacaagaaac tgaaccttct 300
 aaggagttga gtaaactgca gttcagtagt ggtctcaata agtgtgttat acttgctttg 360
 gtgattgcaa tcagcatggg atttggccat ttctatggca c 401

<210> 277
 <211> 401
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(401)
 <223> n = A,T,C or G

<400> 277
 aactttggca acatatctca gcaaaaacta cagctatgtt attcatgcca aaataaaagc 60
 tgtgcagagg agtggctgca atgaggtcac aacgggtggtg gatgtaaaag agatcttcaa 120
 gtctcatca cccatccctc gaactcaagt cccgctcatt acaaattctt cttgccagtg 180
 tccacacatc ctgccccatc aagatgttct catcatgtgt tacgagnggc gctcaaggat 240
 gatgcttctt gaaaattgct tagttgaaaa atggagagat cagcttagta aaagatccat 300
 acagtgggaa gagaggctgc aggaacagcg ganaacagtt caggacaaga agaaaacagc 360
 cgggcgcacc agtcgtagta atcccccaa accaaagggg a 401

<210> 278

<211> 401
 <212> DNA
 <213> Homo sapien
 <220>
 <221> misc_feature
 <222> (1)...(401)
 <223> n = A,T,C or G

<400> 278

aatgagtgtg agaccacaaa tgaatgccgg gaggatgaaa tgtgttgga	60
ggttcctgtt gttatccacg aaatccttgt caagatccct acattctaac accagagAAC	120
cgatgtgttt gccagtcctc aaatgccatg tgccgagAAC tgccccagtc aatagtctac	180
aaatacatga gcatccgacg tgataggtct gtgccatcag acatcttcca gatacaggcc	240
acaactatit atgccaacac catcaatact ttctggatta aatctggaaa tgaaaatgga	300
gagtctacct acgacaacaa anccctgtaa gtgcaatgct tgtgctcgtg aagncattat	360
caggaccaag agaacatatc gtggacctgg agatgctgac a	401

<210> 279
 <211> 401
 <212> DNA
 <213> Homo sapien
 <220>
 <221> misc_feature
 <222> (1)...(401)
 <223> n = A,T,C or G

<400> 279

aaattattgc ctctgataca tacctaagtn aacanaacat taatacctaa gtaaacataa	60
cattacttgg aggggtgcag nttctaantg aaactgtatt tgaaactttt aagtatactt	120
taggaaacaa gcatgaacgg cagtctagaa taccagaaac atctacttgg gtagcttggn	180
gccattatcc tgtggaatct gatatgtctg gnagcatgct attgatggga catgaagaca	240
tctttggaaa tgatgagatt atttcctgtg ttaaaaaaaa aaaaaatctt aaattcctac	300
aatgtgaaac tgaaactaat aattttgatc ctgatgtatg ggacagcgta tctgtaccag	360
gctctaaata acaaaagnta gggngacaag nacatgttcc t	401

<210> 280
 <211> 326
 <212> DNA
 <213> Homo sapien

<400> 280

gaagtggAat tgtataattc aattcgataa ttgatctcat gggctttccc tggaggaaag	60
gttttttttt ttgttttttt ttaagaact tgaaacttgt aaactgagat gtctgtagct	120
tttttgccca tctgtagtgt atgtgaagat ttcaaaacct gagagcactt tttctttgtt	180
tagaattatg agaaaggcac tagatgactt taggatttgc atttttccct ttattgcctc	240
atttcttgtg acgccttggt ggggagggaa atctgtttat tttttcctac aaataaaaaag	300
ctaagattct atatcgcaaa aaaaaa	326

<210> 281
 <211> 374
 <212> DNA
 <213> Homo sapien

<400> 281

caacgcgttt gcaaattatc ccctggtagc ctacttcctt acccccgaat attggttaaga	60	-
tcgagcaatg gcttcaggac atgggttctc ttctcctgtg atcattcaag tgctcactgc	120	
atgaagactg gcttgtctca gtgtttcaac ctcaccaggg ctgtctcttg gtccacacct	180	
cgctccctgt tagtgccgta tgacagcccc catcaaata ccttggccaa gtcacggttt	240	
ctctgtggtc aagggttggt ggctgattgg tggaaagtag ggtggacca aggaggccac	300	-
gtgagcagtc agcaccagtt ctgcaccagc agcgccctcg tctagtggg tgttctgtt	360	
tctctggcc ctgg	374	

<210> 282

<211> 404

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(404)

<223> n = A,T,C or G

<400> 282

agtgtggtgg aattcccga tctanncg cgaactcac aaggcagagt ngccatggag	60
aaaattccag tgcagcatt cttgtcctt gtggccctct cctacactct ggccagagat	120
accacagtca aacctgnagc caaaaaggac acaaaggact ctgcaccaa actgcccan	180
accctctcca gaggttgggg tgaccaactc atctggactc anacatatga agaagctcta	240
tataaatcca agacaagcaa caaaccttg atgattatc atcacttga tgagtgccca	300
cacagtcaag ctttaaagaa agtgtttgct gaaaataaag aaatccagaa attggcagag	360
cagtttgtec tctcaatct ggtttatgaa acaactgaca aaca	404

<210> 283

<211> 184

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(184)

<223> n = A,T,C or G

<400> 283

agtgtggtgg aattcacttg cttaanttgt gggcaaaaga gaaaaaga gattgatcag	60
agcattgtgc aatacagttt cattaactcc ttccctcgct ccccaaaaa tttgaatttt	120
tttttcaaca ctcttacacc tgttatggaa aatgtcaacc tttgtaagaa aacaaaaata	180
aaaa	184

<210> 284

<211> 421

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(421)

<223> n = A,T,C or G

<400> 284

```

ctattaatcc tgccacaata tttttaatta cgtacaaaga tctgacatgt caccagggga    60
cccatttcac ccactgctct gtttggccgc cagtcttttg tctctctctt cagcaatggg    120
gaggcgggata ccctttcctc ggggaanana aatccatggg ttgttgccct tgccaataac    180
aaaaatggtg gaaagtcgag tggcaaagct gttgccattg gcatctttca cgtgaaccac    240
gtcaaaagat ccagggtgcc tctctctgtt ggtgatcaca ccaattcttc ctaggttagc    300
acctccagtc accatacaca ggttaccagt gtcgaacttg atgaaatcag taatcttgcc    360
agtctctaaa tcaatctgaa tggatcatt caccttgatg aggggatcgg ggtagcggat    420
g                                                                    421

```

```

<210> 285
<211> 361
<212> DNA
<213> Homo sapien

```

```

<220>
<221> misc_feature
<222> (1)...(361)
<223> n = A,T,C or G

```

```

<400> 285
ctgggtggta actctttatt tcattgtccg gaanaaagat gggagtggga acagggtgga    60
cactgtgcag gcttcagctt ccactccggg caggattcag gctatctggg accgcaggga    120
ctgccagtg cagagccctg gctcccaggg caggcaggca aggtgacggg actggaagcc    180
cttttcanag ccttggagga gctggtccgt ccacaagcaa tgagtgccac tctgcagttt    240
gcaggggatg gataaacagg gaaacactgt gcattcctca cagccaacag ttaggtctt    300
ggtgaagccc cggcgctgag ctaagctcag gctgttccag ggagccacga aactgcagg    360
a                                                                    361

```

```

<210> 286
<211> 336
<212> DNA
<213> Homo sapien

```

```

<220>
<221> misc_feature
<222> (1)...(336)
<223> n = A,T,C or G

```

```

<400> 286
tttgagtggc agcgccttta tttgtggggg ccttcaaggn agggctcgtg ggggcagcgg    60
ggaggaanag ccganaaact gtgtgaccgg ggcctcaggg ggtgggcatt gggggctcct    120
cttgcanaatg ccattggca tcaccggtgc agccattggg ggcagcgggt accggtcctt    180
tcttgttcaa catagggtag gtggcagcca cgggtccaac tcgcttgagg ctgggccctg    240
ggcgctccat tttgtgttcc angagcatgt ggttctgtgg cgggagcccc acgcaggccc    300
tgaggatgtt ctgatgcag ctgcgctggc ggaaaaa                                                                    336

```

```

<210> 287
<211> 301
<212> DNA
<213> Homo sapien

```

```

<220>
<221> misc_feature
<222> (1)...(301)
<223> n = A,T,C or G

```

<400> 287

tgggtaccaa	atttntttat	ttgaagggaat	ggnacaaatc	aaanaactta	agnggatgtt	60
ttggtacaac	ttatanaaaa	ggnaaaggaa	accccaacat	gcattgcctg	ccttgngac	120
caggaagtc	acccacggc	tatggggaaa	ttancccgag	gcttancctt	cattatcact	180
gtctcccagg	gngngctt	gt	caaaaanata	ttccnccaag	ccaaattcgg	240
nttgcncagg	ttggtcacgt	gggtcaccaa	ttctttgatg	gctttcacct	gctcattcag	300
g						301

<210> 288

<211> 358

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (358)

<223> n = A,T,C or G

<400> 288

aagtttttaa	actttttatt	tgcatattaa	aaaaattgng	cattccaata	attaaaatca	60
tttgaacaaa	aaaaaaaaatg	gcaactctgat	taaactgc	tacagcctgc	aggacacctt	120
gggacagctt	ggttttactc	tanatttcac	tgctgtccca	ccccacttct	tccacccccac	180
ttcttccttc	accaacatgc	aagttcttct	cttccctgcc	agccanata	atagacagat	240
gggaaaggca	ggcgggcct	tcgttgtcag	tagttctttg	atgtgaaagg	ggcagcacag	300
tcatttaaac	ttgatccaac	ctctttgc	cttacaaagt	taaacagcta	aaagaagt	358

<210> 289

<211> 462

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (462)

<223> n = A,T,C or G

<400> 289

ggcatcagaa	atgctgttta	tttctctgct	gtccccaagc	tggctggcct	ttgcagagga	60
gcagacaaca	gatgcatagt	tgggganaaa	gggaggacag	gttccaggat	agagggtgca	120
ggctgaggga	ggaagggtaa	naggaaggaa	ggccatcctg	gatccccaca	tttcagtctc	180
anatgaggac	aaagggactc	ccaagcccc	aatcatcan	aaaacaccaa	ggagcaggag	240
gagcttgagc	aggccccagg	gagcctcana	gccataccag	ccactgtcta	cttcccatcc	300
tcctctccca	ttccctgtct	gtttcanacc	acctccagc	taagccccag	ctccattccc	360
ccaatcctgg	cccttgccag	cttgacagtc	acagtgcctg	gaattccacc	actgaggctt	420
ctcccagttg	gattaggacg	tcgccctgtt	agcatgctgc	cc		462

<210> 290

<211> 481

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (481)

<223> n = A,T,C or G

<400> 290

tacttttcta	aacttttatta	aagaaaaaag	caataagcaa	tggnggtaaa	tctctanaac	60
atacccaatt	ttctgggctt	cctccccga	gaatgtgaca	ttttgatttc	caaacatgcc	120
anaagtgtat	ggttcccaac	tgtactaaag	taggtganaa	gctgaagtcc	tcaagtgttc	180
atcttccaac	ttttcccagt	ctgtggctctg	tctttggatc	agcaataatt	gcctgaacag	240
ctactatggc	ttcgttgatt	tttgtctgta	gctctctgag	ctcctctatg	tgcagcaatc	300
gcanaatttg	agcagcttca	ttaanaactg	catctcctgt	gtcaaaacca	anaatatgtt	360
tgtctaaagc	aacaggtaag	ccctcttttg	tttgatttgc	cttancaact	gcacccctgtg	420
tcaggcgctc	ctgaacccaa	atccgaattg	ccttaagcat	taccaggtaa	tcacatcatgac	480
g						481

<210> 291

<211> 381

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (381)

<223> n = A,T,C or G

<400> 291

tcataagtaat	gtaaaaccat	ttgtttaatt	ctaaatcaaa	tcactttcac	aacagtgtaaa	60
attagtgact	ggttaaggng	tgccactgta	catatcatca	ttttctgact	ggggtcaggga	120
cctggctcta	gtccacaagg	gtggcaggag	gaggggtggag	gctaanaaca	cagaaaacac	180
acaaaanaaa	ggaaagctgc	cttggcanaa	ggatgaggng	gtgagcttgc	cgaaggatgg	240
tgggaagggg	gctccctgtt	ggggccgagc	caggagtccc	aagtcagctc	tcctgcctta	300
cttagctcct	ggcanagggt	gagtggggac	ctacgaggtt	caaaatcaaa	tggcatttgg	360
ccagcctggc	tttactaaca	g				381

<210> 292

<211> 371

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1) ... (371)

<223> n = A,T,C or G

<400> 292

gaaaaaataa	tcggtttaat	tgaaaaacct	gnaggatact	attccactcc	cccanatgag	60
gaggctgagg	anaccaaacc	cctacatcac	ctcgtagcca	cttctgatac	tcttcacgag	120
gcagcaggca	aagacaattc	ccaaaaacctc	nacaaaagca	attccaaggg	ctgctgcagc	180
taccaccanc	acatttttcc	tcagccagcc	cccaatcttc	tccacacagc	cctccttatg	240
gatgccttc	tcgttgaaat	taatcccaca	gcccacagta	acattaatgc	ancaggagtc	300
ggggactcgg	ttcttcgaca	tgggaaggat	tttctcccaa	tctgtgtagt	tagcagcccc	360
acagcactta	a					371

<210> 293

<211> 361

<212> DNA

<213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (361)
 <223> n = A,T,C or G

<400> 293

gatttaaaag	aaaacacttt	attgttcagc	aattaaaagt	tagccaaata	tgtatttttc	60
tccataat	ttt attgngatgt	tatcaacatc	aagtaaaatg	ctcattttca	tcatttgctt	120
ctgttc	catgt tttcttgaac	aegtcttcaa	ttttccttcc	aaaatgctgc	atgccacact	180
tgaggtaacg	aagcanaagt	atTTTTaaac	atgacagcta	anaacattca	tctacagcaa	240
cctatatgct	caatacatgc	cgcgtgatcc	tagtagtttt	ttcacaacct	tctacaagtt	300
tttgga	aaaac atctgttatg	atgactttca	tacaccttca	cctcaaaggc	tttcttgcac	360
c						361

<210> 294
 <211> 391
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (391)
 <223> n = A,T,C or G

<400> 294

tatttttaaag	tttaattatg	attcanaaaa	aatcgagcga	ataactttct	ctgaaaaaat	60
atattgactc	tgtatanacc	acagttattg	gggganaagg	gctggtaggt	taaattatcc	120
tattttttat	tctgaaaatg	atattaatan	aaagtcctcg	ttccagtctg	attataaaga	180
tacatatgcc	caaaatggct	ganaataaat	acaacaggaa	atgcaaaagc	tgtaaagcta	240
agggcatgca	ananaaaatc	tcanaatacc	caaagnggca	acaaggaacg	tttggtgga	300
atttgaagtt	atttcagtca	tctttgtctt	tggtccatg	tttcaggatg	cgtgtgaact	360
cgatgtaatt	gaaattcccc	tttttatcaa	t			391

<210> 295
 <211> 343
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (343)
 <223> n = A,T,C or G

<400> 295

ttcttttgtt	ttattgataa	cagaaactgt	gcataattac	agatttgatg	aggaatctgc	60
aaataataaa	gaatgtgtct	actgccagca	aaatacaatt	attccatgcc	ctctcaacat	120
acaaatatag	agttcttcac	accanattgc	tctggtgtaa	caaagccatt	ttanatgttt	180
aattgtgctt	ctacaaaacc	ttcanagcat	gaggtagtgt	cttttaccta	cnatatatttc	240
cacattttcca	ttattacact	tttagtgagc	taaaatcctt	ttaacatagc	ctgcggatga	300
tctttcacaa	aagccaagcc	tcatttacaa	agggtttatt	tct		343

<210> 296
 <211> 241
 <212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(241)

<223> n = A,T,C or G

<400> 296

```

ttcttgata ttggtgttt ttgtgaaaaa gtttttgttt ttcttctcag tcaactgaat      60
tatttctcta ctttgccctc ctgatgccca catgananaa cttaanataa tttctaacag      120
cttcactttt ggaaaaaaa aaaacctgtt ttctcatgg aaccccagga gttgaaagt      180
gatanatcgc tctcaaatc taaggctctg ttcagcttta cattatgta cctgacgtt      240
t                                                                                   241

```

<210> 297

<211> 391

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(391)

<223> n = A,T,C or G

<400> 297

```

gttgtggctg anaatgctgg agatgctcag ttctctccct cacaaggtag gccacaaatt      60
cttgtgggtg ccctcacatc tggggtcttc aggcaccagc catgctgcc gaggagtgt      120
gtcaggacan accatgtccg tgctaggccc aggcacagcc caaccactcc tcatccaagt      180
ctctcccagg tttctgggtc cgatgggcaa ggatgacccc tccagtggct ggtaccccac      240
catcccacta ccctcacat gctctcactc tccatcaggt cccaatcct ggctccctc      300
ttcacgaact ctcaaagaaa aggaaggata aaacctaat aaaccagaca gaagcagctc      360
tggaagagta caaaaagaca gccagagggtg t                                                                                   391

```

<210> 298

<211> 321

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(321)

<223> n = A,T,C or G

<400> 298

```

caagccaac tgtntccagc tttattaaan atactttcca taaacaatca tggattttca      60
ggcaggacat gggcanacaa tcgttaacag tatacaacaa ctttcaaact cccttnttca      120
atggactacc aaaaatcaaa aagccactat aaaacccaat gaagtcttca tctgatgtc      180
tgaacaggga aagtttaaag ngagggttga catttcacat ttagcatgtt gtttaacaac      240
ttttcacaag cgcacctga ctttcaggaa gtgaaatgaa aatggcanaa tttatctgaa      300
natccacaat ctaaaaatgg a                                                                                   321

```

<210> 299

<211> 401

<212> DNA

<213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(401)
 <223> n = A,T,C or G

<400> 299
 tatcataaag agtgttgaag tttattttatt atagcaccat tgagacattt tgaaattgga 60
 attggtaaaa aaataaaaca aaaagcattt gaattgtatt tggnggaaca gcaaaaaaag 120
 agaagtatca tttttctttg tcaaattata ctgtttccaa acatttttga aataaataac 180
 tggaaatttg tcggtcactt gcaactgggtg acaagattag aacaagagga acacatatgg 240
 agttaaatat tttttgttgg gatttcanat agagtttggg ttataaaaag caaacagggc 300
 caacgtccac accaaattct tgatcaggac caccaatgtc atagggngca atatctacaa 360
 taggtagtct cacagccttg cgtgttcgat attcaaagac t 401

<210> 300
 <211> 188
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(188)
 <223> n = A,T,C or G

<400> 300
 tgaatgcttt gtcataattaa gaaagttaaa gtgcaataat gtttgaanac aataagtggg 60
 ggtgtatctt gtttctaata agataaactt ttttgtcttt gctttatctt attagggagt 120
 tgtatgtcag tgtataaaac atactgtgtg gtataacagg cttataaat tctttaaaag 180
 gaaaaaaa 188

<210> 301
 <211> 291
 <212> DNA
 <213> Homo sapien

<400> 301
 aagattttgt tttattttat tatggctaga aagacactgt tatagccaaa atcggaatg 60
 aactaaaga aatcctctgt gcttttcaat atgcaaatat atttcttcca agagttgccc 120
 tgggtgact tcaagagttc atgttaactt cttttctgga aacttcttt tcttagttgt 180
 tgtattcttg aagagcctgg gccatgaaga gcttgccata gttttgggca gtgaactcct 240
 tgatgttctg gcagtaagtg tttatctggc ctgcaatgag cagcgagtc a 291

<210> 302
 <211> 341
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(341)
 <223> n = A,T,C or G

<400> 302
 tgatttttca taattttatt aatnatcac tgggaaaact aatgggtcgc gtatcacaca 60


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attacactac aatctgatag gagtggtaaa accagccaat ggaatccagg taaagtacaa 120
aaacgccacc ttttattgtc ctgtcttatt tctcgggaag gagggttcta ctttacacat 180
ttcatgagcc agcagtggaac ttgagttaca atgtgtaggt tccttgtggt tatagctgca 240
gaagaagcca tcaaattctt gaggacttga catctctcgg aaagaagcaa actagtggat 300
ccccggggt gcaggaattc gatatcaagc ttatcgatac c 341

```

<210> 303

<211> 361

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(361)

<223> n = A,T,C or G

<400> 303

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tgcagacagt aaatnaattt tatttgngtt cacagaacat actaggcgat ctgcacagtc 60
gctcgtgac agcccaccaa cccccaaccc tntacctgc agccacccta aaggcgactt 120
caanaanatg gaaggatctc acggatctca ttcctaattg tccgccgaag tctcacacag 180
tanacagacg gagttganat gctggaggat gcagtcacct cctaaactta cgaccacca 240
ccanaettca tcccagccgg gacgtctctc cccaccggag tcctcctccat ttcttctcct 300
actttgccgc agttccaggn gtctgtcttc caccagtccc acaaagctca ataaatacca 360
a 361

```

<210> 304

<211> 301

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(301)

<223> n = A,T,C or G

<400> 304

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ctctttacaa cagcctttat ttncggccct tgatcctgct cggatgctgg tggaggccct 60
tagctccgcc cgccaggctc tgtgccgctt ccccgaggc gcanattcat gaacacgggtg 120
ctcagggggt tgaggccgta ctccccagc gggagctggt cctccagggg cttccctcgg 180
aaggtcagcc anaacaggtc gtctgcaca ccctccagcc cgctcacttg ctgcttcagg 240
tgggccacgg tctgcgtcag ccgcacctcg taggtgctgc tgcggccctt gttattcctc 300
a 301

```

<210> 305

<211> 331

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(331)

<223> n = A,T,C or G

<400> 305

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ganaggctag taacatcagt tttattgggt tggggnggca accatagcct ggctgggggn 60

```

ggggctggcc	ctcacaggtt	gttgagttcc	agcagggtct	ggccaaggt	ctggtgaatc	120
tcgacgttct	cctccttggc	actggccaag	gtctcttcta	ggcatcgat	ggttttctcc	180
aactttgcca	canacctctc	ggcaaacctc	gtcgggtct	cancctcctt	cagcttctcc	240
tccaacagtt	tgatctctc	ttcatattta	tcttcttggg	gggaatactc	ctcctctgag	300
gccatcaggg	acttgagggc	ctggtccatg	g			331

<210> 306

<211> 457

<212> DNA

<213> Homo sapien

<400> 306

aatatgtaaa	ggtaataact	tttattatat	ttaaagacaat	gcaaacgaaa	aacagaattg	60
agcagtgc	aatTTaaagg	actgttttgt	tctcaaagtt	gcaagtttca	aagccaaaag	120
aattatatgt	atcaaatata	taagtaaaaa	aaagtttagac	tttcaagcct	gtaatcccag	180
cactttggga	ggctgaggca	ggtggatcac	taacattaaa	aagacaacat	tagattttgt	240
cgatttatag	caattttata	aatatataac	tttgcactt	ggatcctgaa	gcaaaaataat	300
aaagtgaatt	tgggattttt	gtacttggtg	aaaagtttaa	caccctaaat	tcacaactag	360
tggatcccc	gggctgcagg	aattcgatat	caagcttata	gataccgtcg	acctcgaggg	420
ggggcccggt	acccaattcg	ccctatagtg	agtcgta			457

<210> 307

<211> 491

<212> DNA

<213> Homo sapien

<400> 307

gtgcttgga	ggaaccggc	gtcgttccc	caccccgcc	ggccgccc	agccagccct	60
ccgtcacctc	ttcacgcac	cctcggactg	ccccaggcc	cccgccgcg	ctccagcgcc	120
gcgcagccac	cgccgccc	gccgcctctc	cttagtcgc	gccatgacga	ccgcgtccac	180
ctcgcagggtg	cgccagaact	accaccagga	ctcagaggcc	gccatcaacc	gccagatcaa	240
cctggagctc	tacgcctcct	acgtttacct	gtccatgtct	tactactttg	accgcgatga	300
tgtggctttg	aagaactttg	ccaaataactt	tcttaccac	tctcatgagg	agaggggaaca	360
tgctgagaaa	ctgatgaagc	tgacgaacca	acgaggtggc	cgaatcttcc	ttcaggatat	420
caagaaacca	gactgtgatg	actgggagag	cgggctgaat	gcaatggagt	gtgcattaca	480
tttgga	aaaaa	a				491

<210> 308

<211> 421

<212> DNA

<213> Homo sapien

<400> 308

ctcagcgctt	cttctttctt	ggtttgatcc	tgactgctgt	catggcgtgc	cctctggaga	60
aggccctgga	tgtgatggtg	tccaccttcc	acaagtactc	gggcaaagag	ggtgacaagt	120
tcaagctcaa	caagtcagaa	ctaaaggagc	tgctgaccog	ggagctgccc	agcttcttgg	180
ggaaaaggac	agatgaagct	gctttccaga	agctgatgag	caacttggac	agcaacaggg	240
acaacyaggt	ggacttccaa	gagtactgtg	tcttctgtc	ctgcatcgcc	atgatgtgta	300
acgaattctt	tgaaggcttc	ccagataagc	agcccaggaa	gaaatgaaaa	ctcctctgat	360
gtggttgggg	ggtctgccag	ctggggccct	ccctgtcgcc	agtgggcact	tttttttttc	420
c						421

<210> 309

<211> 321

<212> DNA

<213> Homo sapien

<400> 309

accaaagggc	ggatgacgcc	ggtgcagcgg	gggggcccgg	gggccctggg	ggccctggga	60
tggggaaccg	cggtggttc	cgcggaggtt	tcggcagtg	catccggggc	cggggtcgcg	120
gccgtggacg	gggccggggc	caggcccgcg	gagctcgcg	aggcaaggcc	gaggataagg	180
agtggatgcc	cgtcaccaag	ttgggccgct	tggtcaagga	catgaagatc	aagtcctgg	240
aggagatcta	tctcttctcc	ctgccatta	aggaatcaga	gatcattgat	ttcttcctgg	300
gggcctctct	caaggatgag	g				321

<210> 310

<211> 381

<212> DNA

<213> Homo sapien

<400> 310

ttaaccagcc	atattggctc	aataaatagc	ttcggtaagg	agttaatttc	cttctagaaa	60
tcagtgccta	tttttctgg	aaactcaatt	ttaaatagtc	caattccatc	tgaagccaag	120
ctgttgcat	tttcattcgg	tgacattctc	tcccatgaca	cccagaagg	gcagaagaac	180
cacatttttc	atttatagat	gtttgcattc	tttgatttaa	aattattttg	aaggggttgc	240
ctcattggat	ggctttttt	tttttctcc	agggagaagg	ggagaaatgt	acttggaaat	300
taatgtatgt	ttacatctct	ttgcaaattc	ctgtacatag	agatatattt	tttaagtgtg	360
aatgtaacaa	catactgtga	a				381

<210> 311

<211> 538

<212> DNA

<213> Homo sapien

<400> 311

tttgaattta	caccaagaac	ttctcaataa	aagaaaatca	tgaatgctcc	acaattttcaa	60
cataccacaa	gagaagttaa	tttcttaaca	ttgtgttcta	tgattatttg	taagaccttc	120
accaagtctc	gatatctttt	aaagacatag	ttcaaaattg	cttttgaaaa	tctgtattct	180
tgaataatc	cttgttgtgt	attaggtttt	taaataaccag	ctaaaggatt	acctcactga	240
gtcatcagta	ccctcctatt	cagctcccca	agatgatgtg	tttttgctta	ccctaagaga	300
ggttttcttc	ttatttttag	ataattcaag	tgcttagata	aattatgttt	tctttaagtg	360
tttatggtaa	actcttttaa	agaaaattta	atatgttata	gctgaatctt	tttggttaact	420
ttaaatcttt	atcatagact	ctgtacatat	gttcaaatta	gctgcttgcc	tgatgtgtgt	480
atcatcggtg	ggatgacaga	acaaacatat	ttatgatcat	gaataatgtg	ctttgtaa	538

<210> 312

<211> 176

<212> DNA

<213> Homo sapien

<400> 312

ggaggagcag	ctgagagata	gggtcagtga	atgcggttca	gcctgctacc	tctcctgtct	60
tcatagaacc	attgccttag	aattattgta	tgacacgttt	ttgttggtt	aagctgtaag	120
gtttgttct	ttgtgaacat	gggtattttg	aggggaggg	ggaggagta	gggaag	176

<210> 313

<211> 396

<212> DNA

<213> Homo sapien

<400> 313

ccagcacc	ccc	caggccctgg	gggacctggg	ttctcagact	gccaaagaag	ccttgccatc	60
tggegetccc	atggetcttg	caacatctcc	ccttcgtttt	tgaggggggc	atgccggggg		120
agccaccagc	ccctcactgg	gttcggagga	gagtcaggaa	gggccaaagca	cgacaaagca		180
gaaacatcgg	atttggggaa	cgcggtgtcaa	tcccttggtc	cgccagggtg	ggcgggagag		240
actgttctgt	tccttggtga	actgtgttgc	tgaaagacta	cctcgttctt	gtcttgatgt		300
gtcacccggg	caactgcctg	ggggcgggga	tgggggcagg	gtggaagcgg	ctccccattt		360
tataccaaag	gtgctacatc	tatgtgatgg	gtgggg				396

<210> 314

<211> 311

<212> DNA

<213> Homo sapien

<400> 314

cctcaacatc	ctcagagagg	actggaagcc	agtccttacg	ataaactcca	taatttatgg	60
cctgcagtat	ctcttcttgg	agcccaaccc	cgaggaccca	ctgaacaagg	aggccgcaga	120
ggctctgcag	aacaaccggc	ggctgtttga	gcagaacgtg	cagcgctcca	tgccgggtgg	180
ctacatcggc	tccacctact	ttgagcgctg	cctgaaatag	ggttggcgca	taccaccccc	240
cgccacggcc	acaagccctg	gcatccccctg	caaatatatta	ttggggggcca	tgggtagggg	300
tttggggggc	g					311

<210> 315

<211> 336

<212> DNA

<213> Homo sapien

<400> 315

ttttagaacat	ggttatcatc	caagactact	ctaccctgca	acattgaact	cccaagagca	60
aatccacatt	cctcttgagt	tctgcagctt	ctgtgtaaat	agggcagctg	tcgtctatgc	120
cgtagaatca	catgatctga	ggaccattca	tggaaagctg	taaatagcct	agtctggggg	180
gtcttccata	aagttttgca	tggagcaaac	aaacaggatt	aaactaggtt	tggttccttc	240
agccctctaa	aagcataggg	cttagcctgc	aggcttcctt	gggctttctc	tgtgtgtgta	300
gttttgtaaa	cactatagca	tctgttaaga	tccagt			336

<210> 316

<211> 436

<212> DNA

<213> Homo sapien

<400> 316

aacatgggtct	gcgtgcctta	agagagacgc	ttcctgcaga	acaggacctg	actacaaaga	60
atgtttccat	tggaattggt	ggtaaagact	tggagtttac	aatctatgat	gatgatgatg	120
tgtctccatt	cctggaaggt	cttgaagaaa	gaccacagag	aaaggcacag	cctgctcaac	180
ctgctgatga	acctgcagaa	aaggctgatg	aaccaatgga	acattaagtg	ataagccagt	240
ctatatatgt	attatcaaatt	atgtaagaat	acaggcacca	catactgatg	acaataatct	300
atactttgaa	ccaaaagttg	cagagtgggtg	gaatgctatg	ttttaggaat	cagtccagat	360
gtgagttttt	tccaagcaac	ctcactgaaa	cctatataat	ggaatacatt	tttctttgaa	420
aggggtctgta	taatca					436

<210> 317

<211> 196

<212> DNA

<213> Homo sapien

<400> 317

tattccttgt gaagatgata tactatTTTT gttaagcgtg tctgtattta tgggtgagga	60
gctgctggct tgcagtgcgc gtgcacgtgg agagctgggt cccggagatt ggacggcctg	120
atgctccctc ccctgccctg gtccagggaa gctggccgag ggtcctggct cctgaggggc	180
atctgccctt ccccca	196

<210> 318

<211> 381

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(381)

<223> n = A,T,C or G

<400> 318

gacgcttmg ccgtaacgat gatcggagac atcctgctgt tggggacgtt gctgatgaat	60
gcggggcggg tgctgaactt taagctgaaa aagaaggaca cncagggcct tggggaggag	120
tncagggagc ccaacacagg tgacaacatc cgggaattct tgctgancct cagatacttt	180
cnaatcttca tcnccctgtg gaacatcttc atgatgttct gcatgattgt gctgntcggc	240
tcttgaatcc cancgatgaa accannaact cactttcccg ggatgccgan tctccattcc	300
tccattcctg atgacttcaa naatgttttt gaccaaaaaa ccgacaacct tcccagaaag	360
tccaagctcg tgggtggngg a	381

<210> 319

<211> 506

<212> DNA

<213> Homo sapien

<400> 319

ctaagcttta cgaatggggg gacaacttat gataaaaact agagctagtg aattagccta	60
tttgtaaata cttttgttat aattgatagg atacatcttg gacatggaat tgtaagcca	120
cctctgagca gtgtatgtca ggacttggtc attagggttg cagcagaggg gcagaaggaa	180
ttatacaggt agagatgtat gcagatgtgt ccataatgt ccataattac attttgatag	240
ccattgatgt atgcatctct tggctgtact ataagaacac attaatcaa tggaaataca	300
ctttgctaatt attttaattg tatagatctg ctaatgaatt ctcttaaaaa catactgtat	360
tctgttctg tgtgtttcat tttaaattga gcattaaggg aatgcagcat ttaaatacaga	420
actctgcaa tgcttttatc tagaggcgtg ttgccatttt tgtcttatat gaaatttctg	480
tccaagaaa ggcaggatta catctt	506

<210> 320

<211> 351

<212> DNA

<213> Homo sapien

<400> 320

ctgacctgca ggacgaaacc atgaagagcc tgatccttct tgccatcctg gccgccttag	60
cggtagtaac tttgtgttat gaatcacatg aaagcatgga atcttatgaa cttaatccct	120
tcatcaacag gagaaatgca aataccttca tatccctca gcagagatgg agagctaaag	180
tccaagagag gatccgagaa cgctctaagc ctgtccacga gctcaatagg gaagcctgtg	240
atgactacag actttgcgaa cgctacgcca tggtttatgg atacaatgct gcctataatc	300
gctacttcag gaagcgccga gggaccaaat gagactgagg gaagaaaaa a	351

<210> 321

<211> 421
 <212> DNA
 <213> Homo sapien

<400> 321

ctcggaggcg	ttcagctgct	tcaagatgaa	gctgaacatc	tccttcccag	ccactggctg	60
ccagaaactc	attgaagtgg	acgatgaacg	caaacttcgt	actttctatg	agaagcgtat	120
ggccacagaa	gttgctgctg	acgctctggg	tgaagaatgg	aagggttatg	tggtccgaat	180
cagtgggtggg	aacgacaaac	aaggtttccc	catgaagcag	ggtgtcttga	cccatggccg	240
tgtccgcctg	ctactgagta	aggggcattc	ctgttacaga	ccaaggagaa	ctggagaaaag	300
aaagagaaaa	tcagttcgtg	gttgcatattg	ggatgcaa	ctgagcgttc	tcaacttggg	360
tattgtaaaa	aaaggagaga	aggatattcc	tggactgact	gatactacag	tgccctcgccg	420
c						421

<210> 322
 <211> 521
 <212> DNA
 <213> Homo sapien

<400> 322

agcagctctc	ctgccacagc	tcctcacccc	ctgaaaatgt	tcgcctgctc	caagtttgtc	60
tccactccct	ccttggtcaa	gagcacctca	cagctgctga	gccgtccgct	atctgcagtg	120
gtgctgaaac	gaccggagat	actgacagat	gagagcctca	gcagcttggc	agtctcatgt	180
ccccttacct	cacttgtctc	tagccgcagc	ttccaaacca	gcgccatttc	aaggacatc	240
gacacagcag	ccaagttcat	tggagctggg	gctgccacag	ttgggggtggc	tggttctggg	300
gctgggattg	gaactgtgtt	tgggagcctc	atcattgggt	atgccaggaa	cccttctctg	360
aagcaacagc	tcttctocta	cgccattctg	ggctttgccc	tctcgagggc	catggggctc	420
ttttgtctga	tggtagcctt	tctcatcctc	tttgccatgt	gaaggagccg	tctccacctc	480
ccatagttct	cccgcgtctg	gttggtcccg	tgtgttcctt	t		521

<210> 323
 <211> 435
 <212> DNA
 <213> Homo sapien

<400> 323

ccgaggctgc	acgcgtgaga	cttctccgcc	gcagacgccg	ccgcgatggc	ctacgtcgcc	60
tctacactgc	tggtgcccct	agggggcaac	tcctccccca	gcgccaagga	catcaagaag	120
atcttgga	gcgtgggtat	cgaggcggac	gacgaccggc	tcaacaaggt	tatcagtga	180
ctgaatggaa	aaaacattga	agacgtcatt	gcccagggta	ttggcaagct	tgccagtgt	240
cctgctgggtg	gggctgtagc	ogtctctgct	gccccaggct	ctgcagcccc	tgctgctggg	300
tctgcccctg	ctgcagcaga	ggagaagaaa	gatgagaaga	aggaggagtc	tgaagagtca	360
gatgatgaca	tgggatttgg	cctttttgat	taaattcctg	ctcccctgca	aataaagcct	420
ttttacacat	ctcaa					435

<210> 324
 <211> 521
 <212> DNA
 <213> Homo sapien

<400> 324

aggagatcga	ctttcggtgc	ccgcaagacc	agggctggaa	cgccgagatc	acgctgcaga	60
tggtgcagta	caagaatcgt	caggccatcc	tggcggtcaa	atccacgcgg	cagaagcagc	120
agcacctggg	ccagcagcag	ccccctctgc	agccgcagcc	gcagccgcag	ctccagcccc	180
aacccagcc	tcagcctcag	ccgcaacccc	agccccaatc	acaacccag	cctcagcccc	240

```

aaccceaagcc tcagccccag cagctccacc cgtatccgca tccacatcca catccacact    300
ctcctectca ctgcaccca caccctcacc cgcacccgca tccgcaccaa ataccgcacc    360
cacacccaca gccgcactcg cagcgcacg ggcacccggt tctccgcagc acctccaaact    420
ctgctgaaa ggggcagctc ccgggcaaga caaggttttg aggacttgag gaagtgggac    480
gagcacattt ctattgtctt cacttggatc aaaagcaaaa c                    521

```

<210> 325

<211> 451

<212> DNA

<213> Homo sapien

<400> 325

```

attttcattt ccattaacct ggaagctttc atgaatattc tcttctttta aaacatttta    60
acattattta aacagaaaaa gatgggctct ttctggttag ttgttcatg atagcagaga    120
tatttttact tagattactt tgggaatgag agattgttgt cttgaactct ggcactgtac    180
agtgaatgtg tctgtagttg tggttagttg cattaagcat gtataacatt caagtatgtc    240
atccaaataa gaggcataata cattgaattg tttttaatcc tctgacaagt tgactcttcg    300
acccccaccc ccaccaaga cattttaata gtaaatagag agagagagaa gagttaatga    360
acatgaggtg gtgttcact ggcaggatga cttttcaata gtcacaaatc atttcagtgc    420
ctttatcact tgaattatta acttaatttg a                    451

```

<210> 326

<211> 421

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(421)

<223> n = A,T,C or G

<400> 326

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cgcggtcgta agggctgagg atttttggtc cgcacgctcc tgctcctgac tcaccgctgt    60
tcgctctcgc cgaggaacaa gtcggtcagg aagcccgccg gcaacagcca tggcttttaa    120
ggataccgga aaaacacccg tggagccgga ggtggcaatt caccgaattc gaatcacctt    180
aacaagccgc aacgtaaaat ccttggaaaa ggtgtgtgct gacttgataa gaggcgcaaa    240
agaaaagaat ctcaaagtga aaggaccagt tcgaatgcct accaagactt tgagantcac    300
tacaagaaaa actccttggt gtgaagggtc taagacgtgg gatcgtttcc agatgagaat    360
tcacaagcga ctattgact tgcacagtcc ttctgagatt gttaagcaga ttacttccat    420
c                    421

```

<210> 327

<211> 456

<212> DNA

<213> Homo sapien

<400> 327

```

atcttgacga ggctgcggtg tctgctgcta ttctccgagc ttcgcaatgc cgcctaagga    60
cgacaagaag aagaaggacg ctggaaagtc ggccaagaaa gacaaagacc cagtgaacaa    120
atccgggggc aaggccaaaa agaagaagtg gtccaaaggc aaagttcggg acaagctcaa    180
taacttagtc ttgtttgaca aagctaccta tgataaactc tgtaaggaag ttcccaacta    240
taaacttata accccagctg tggctctctg gagactgaag attcgaggct ccctggccag    300
ggcagccctt caggagctcc ttagtaaaag acttatcaaa ctggtttcaa agcacagagc    360
tcaagtaatt tacaccagaa ataccaaggg tggagatgct ccagctgctg gtgaagatgc    420
atgaataggt ccaaccagct gtacatttgg aaaaat                    456

```

<210> 328
 <211> 471
 <212> DNA
 <213> Homo sapien

```

<400> 328
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caggaagac agggcgacct ggaagtccaa ctacttcctt aagatcatcc aactattgga      120
tgattatccg aaatgtttca ttgtgggagc agacaatgtg ggctccaagc agatgcagca      180
gatccgcatg tcccttcgcg ggaaggctgt ggtgctgatg ggcaagaaca ccatgatgcg      240
caaggccatc cgagggcacc tggaaaacaa cccagctctg gagaaactgc tgcctcatat      300
ccgggggaat gtgggctttg tgttcaccaa ggaggacctc actgagatca gggacatgtt      360
gctggccaat aaggtgccag ctgctgcccg tgctggtgcc attgccccat gtgaagtcac      420
tgtgcagacc cagaacactg gtctcggggc cgagaagacc tcctttttcc a              471

```

<210> 329
 <211> 278
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1) ... (278)
 <223> n = A,T,C or G

```

<400> 329
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aaattgagat gccccccag gccagcaaat gttccttttt gttcaaagtc tattttttatt      120
ccttgatatt tttctttttt tttttttttt ttgnnggatg ggacttgtga atttttctaa      180
aggtgctatt taacatggga gganagcgtg tgcggctcca gcccagcccg ctgctcactt      240
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```

<210> 330
 <211> 338
 <212> DNA
 <213> Homo sapien

```

<400> 330
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cacaacatt attataataa acaccctcac cactacaatc ttcctaggaa caacatatga      120
cgcactctcc cctgaactct acacaacata ttttgtcacc aagaccctac ttctaacctc      180
cctgttctta tgaattcgaa cagcataccc cggattccgc tacgaccaac tcatacacct      240
cctatgaaaa aacttcctac cactcacctt agcattactt atatgatatg tctccatacc      300
cattacaatc tccagcatte cccctcaaac ctaaaaaa              338

```

<210> 331
 <211> 2820
 <212> DNA
 <213> Homo sapiens

```

<400> 331
tggcaaaatc ctggagccag aagaaaggac agcagcattg atcaatctta cagctaacat 60
gttgtagctg gaaaacaatg cccagactca atttagtgag ccacagtaca cgaacctggg 120

```



```

gtcctgaac agcatggacc agcagattcg gaacggctcc tcgtccacca gtccctataa 180
cacagaccac ggcgagaaca ggcgtcacggc gccctcgccc tacgcacagc ccagccccac 240
cttcgatgct ctctctccat ccccgccat cccctccaac accgactacc caggcccgca 300
cagttccgac gtgtccttcc agcagtcgag caccgccaag tcggccacct ggacgtattc 360
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gatgaccca cctcctcagg gagctgttat ccgcgccatg cctgtctaca aaaaagctga 480
gcacgtcacg gaggtggtga agcgggtgcc caaccatgag ctgagccgtg agttcaacga 540
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ctaaatttca ctactagatt gactaactca aatacacatt tgctactgtt gtaagaattc 2820

```

<210> 332

<211> 2270

<212> DNA

<213> Homo sapiens

<400> 332

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tcgttgatat caaagacagt tgaaggaaat gaattttgaa acttcacggt gtgccaccct 60
acagtactgc cctgaccctt acatccagcg tttcgtagaa acccagctca tttctcttgg 120

```

```

aaagaaagtt attaccgatc caccatgtcc cagagcacac agacaaatga attcctcagt 180
ccagagggttt tccagcatat ctgggatttt ctggaacagc ctatatgttc agttcagccc 240
attgacttga actttgtgga tgaaccatca gaagatggtg cgacaaacaa gattgagatt 300
agcatggact gtatccgcat gcaggactcg gacctgagtg accccatgtg gccacagtac 360
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tggacgtatt ccactgaact gaagaaactc tactgccaaa ttgcaaagac atgccccatc 660
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```

<210> 333

<211> 2816

<212> DNA

<213> Homo sapiens

<400> 333

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aaagaaagtt attaccgatc caccatgtcc cagagcacac agacaaatga attcctcagt 180
ccagagggttt tccagcatat ctgggatttt ctggaacagc ctatatgttc agttcagccc 240
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agtccctata acacagacca cgcgcagaac agcgtcacgg cgccctcgcc ctacgcacag 480
cccagctcca ccttcgatgc tctctctcca tcaccgcgca tcccctccaa caccgactac 540
ccaggcccgc acagtttcga cgtgtccttc cagcagtcga gcaccgcca gtctggccacc 600
tggacgtatt ccactgaact gaagaaactc tactgccaaa ttgcaaagac atgccccatc 660

```

```

cagatcaagg tgatgacccc acctcctcag ggagctgtta tccgcgccat gcctgtctac 720
aaaaaagctg agcacgtcac ggaggtgggt aagcgggtgcc ccaaccatga gctgagccgt 780
gaattcaacg agggacagat tgccccctct agtcatttga ttcgagtaga ggggaacagc 840
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<210> 334

<211> 2082

<212> DNA

<213> Homo sapiens

<400> 334

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tgtaacacag tggtaagtct ttgtgtatct aaacatagct aaacaccaa aggtatagta 180
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gcacagccca gctccacctt cgatgctctc tctccatcac ccgccatccc ctccaacacc 660

```

```

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<210> 335

<211> 4849

<212> DNA

<213> Homo sapiens

<400> 335

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aaagaaagtt attaccgatc caccatgtcc cagagcacac agacaaatga attcctcagt 180
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aatccccata	aataacagta	tgtgggatat	tgaatgttaa	agggatattt	ttttcttatt	4620
atttttataa	ttgtacaaaa	ttaagcaaat	gttaaaagt	ttatatgctt	tattaatggt	4680

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ttcaaaagggt attatacatg tgatacattt ttttaagcttc agttgcttgt cttctggtac 4740
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gacatgcaat aaaattttaa aaataaataa aaactaatta agaaataaa 4849

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<210> 336

<211> 1386

<212> DNA

<213> Homo sapiens

<400> 336

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aacacagacc acgcgcagaa cagcgtcagc gcgccctcgc cctacgcaca gccagctcc 180
accttcgatg ctctctctcc atcacccgcc atcccctcca acaccgacta cccaggcccg 240
cacagtttcg acgtgtcctt ccagcagtcg agcaccgcca agtcggccac ctggacgtat 300
tccactgaac tgaagaaact ctactgccaa attgcaaaga catgccccat ccagatcaag 360
gtgatgaccc cacctcctca gggagctgtt atccgcgcca tgctgtcta caaaaaagct 420
gagcacgtca cggaggtggt gaagcgggtgc cccaaccatg agctgagccg tgaattcaac 480
gagggacaga ttgccctcc tagtcatttg attcgagtag aggggaacag ccatgcccag 540
tatgtagaag atcccatcac aggaagacag agtgtgctgg taccttatga gccacccag 600
gttggcactg aattcacgac agtcttgtag aatttcattg gtaacagcag ttgtgttggg 660
gggatgaacc gccgtccaat ttaatacatt gttactctgg aaaccagaga tgggcaagtc 720
ctggggccgac gctgctttga ggcccggatc tgtgcttgcc caggaagaga caggaaggcg 780
gatgaagata gcatcagaaa gcagcaagtt tcggacagta caaagaacgg tgatggtacg 840
aagcgcccggt ttcgtcagaa cacacatggt atccagatga catccatcaa gaaacgaaga 900
tccccagatg atgaactgtt atacttacca gtgaggggccc gtgagactta tgaaatgctg 960
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attcctgatg gcatgggagc caacattccc atgatgggca cccacatgcc aatggctgga 1260
gacatgaatg gactcagccc caccaggcca ctccctcccc cactctccat gccatccacc 1320
tcccactgca cccccacc tccgtatccc acagattgca gcattgtcag gatctggcaa 1380
gtctga 1386

```

<210> 337

<211> 1551

<212> DNA

<213> Homo sapiens

<400> 337

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gattttctgg aacagcctat atgttcagtt cagccatttg acttgaactt tgtggatgaa 120
ccatcagaag atggtgcgac aaacaagatt gagattagca tggactgtat ccgcatgcag 180
gactcggacc tgagtgcacc catgtggcca cagtacacga acctggggct cctgaacagc 240
atggaccagc agattcagaa cggctcctcg tccaccagtc cctataacac agaccacgcg 300
cagaacagcg tcacggcgcc ctgcacctac gcacagccca gctccactt cgatgctctc 360
tctccatcac ccgccatccc ctccaacacc gactaccag gccgcacag ttctgacgtg 420
tccttcagc agtcgagcac cgccaagtgc gccacctgga cgtattccac tgaactgaag 480
aaactctact gccaaattgc aaagacatgc cccatccaga tcaaggtgat gacccacct 540
cctcaggag ctgttatccg cgccatgcct gtctacaaaa aagctgagca cgtcacggag 600
gtggtgaagc ggtgccccaa ccatgagctg agccgtgaat tcaacgagg acagattgcc 660
cctcctagtc atttgattcg agtagagggg aacagccatg cccagtatgt agaagatccc 720

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atcacaggaa gacagagtgt gctggtacct tatgagccac cccaggttgg cactgaattc 780
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ccaattttaa tcattgttac tctggaaacc agagatgggc aagtcctggg ccgacgctgc 900
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agccccaccc aggcactccc tccccactc tccatgccat ccacctccca ctgcacaccc 1500
ccacctcctg atccacaga ttgcagcatt gtcaggatct ggcaagtctg a 1551

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<210> 338

<211> 586

<212> PRT

<213> Homo sapiens

<400> 338

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Met Leu Tyr Leu Glu Asn Asn Ala Gln Thr Gln Phe Ser Glu Pro Gln
      5                      10                      15

```

```

Tyr Thr Asn Leu Gly Leu Leu Asn Ser Met Asp Gln Gln Ile Arg Asn
      20                      25                      30

```

```

Gly Ser Ser Ser Thr Ser Pro Tyr Asn Thr Asp His Ala Gln Asn Ser
      35                      40                      45

```

```

Val Thr Ala Pro Ser Pro Tyr Ala Gln Pro Ser Pro Thr Phe Asp Ala
      50                      55                      60

```

```

Leu Ser Pro Ser Pro Ala Ile Pro Ser Asn Thr Asp Tyr Pro Gly Pro
      65                      70                      75                      80

```

```

His Ser Ser Asp Val Ser Phe Gln Gln Ser Ser Thr Ala Lys Ser Ala
      85                      90                      95

```

```

Thr Trp Thr Tyr Ser Thr Glu Leu Lys Lys Leu Tyr Cys Gln Ile Ala
      100                      105                      110

```

```

Lys Thr Cys Pro Ile Gln Ile Lys Val Met Thr Pro Pro Pro Gln Gly
      115                      120                      125

```

```

Ala Val Ile Arg Ala Met Pro Val Tyr Lys Lys Ala Glu His Val Thr
      130                      135                      140

```

```

Glu Val Val Lys Arg Cys Pro Asn His Glu Leu Ser Arg Glu Phe Asn
      145                      150                      155                      160

```

```

Glu Gly Gln Ile Ala Pro Pro Ser His Leu Ile Arg Val Glu Gly Asn
      165                      170                      175

```

```

Ser His Ala Gln Tyr Val Glu Asp Pro Ile Thr Gly Arg Gln Ser Val

```

180	185	190
Leu Val Pro Tyr Glu Pro Pro Gln Val Gly Thr Glu Phe Thr Thr Val		
195	200	205
Leu Tyr Asn Phe Met Cys Asn Ser Ser Cys Val Gly Gly Met Asn Arg		
210	215	220
Arg Pro Ile Leu Ile Ile Val Thr Leu Glu Thr Arg Asp Gly Gln Val		
225	230	235
Leu Gly Arg Arg Cys Phe Glu Ala Arg Ile Cys Ala Cys Pro Gly Arg		
245	250	255
Asp Arg Lys Ala Asp Glu Asp Ser Ile Arg Lys Gln Gln Val Ser Asp		
260	265	270
Ser Thr Lys Asn Gly Asp Gly Thr Lys Arg Pro Phe Arg Gln Asn Thr		
275	280	285
His Gly Ile Gln Met Thr Ser Ile Lys Lys Arg Arg Ser Pro Asp Asp		
290	295	300
Glu Leu Leu Tyr Leu Pro Val Arg Gly Arg Glu Thr Tyr Glu Met Leu		
305	310	315
Leu Lys Ile Lys Glu Ser Leu Glu Leu Met Gln Tyr Leu Pro Gln His		
325	330	335
Thr Ile Glu Thr Tyr Arg Gln Gln Gln Gln Gln His Gln His Leu		
340	345	350
Leu Gln Lys Gln Thr Ser Ile Gln Ser Pro Ser Ser Tyr Gly Asn Ser		
355	360	365
Ser Pro Pro Leu Asn Lys Met Asn Ser Met Asn Lys Leu Pro Ser Val		
370	375	380
Ser Gln Leu Ile Asn Pro Gln Gln Arg Asn Ala Leu Thr Pro Thr Thr		
385	390	395
Ile Pro Asp Gly Met Gly Ala Asn Ile Pro Met Met Gly Thr His Met		
405	410	415
Pro Met Ala Gly Asp Met Asn Gly Leu Ser Pro Thr Gln Ala Leu Pro		
420	425	430
Pro Pro Leu Ser Met Pro Ser Thr Ser His Cys Thr Pro Pro Pro Pro		
435	440	445
Tyr Pro Thr Asp Cys Ser Ile Val Ser Phe Leu Ala Arg Leu Gly Cys		
450	455	460
Ser Ser Cys Leu Asp Tyr Phe Thr Thr Gln Gly Leu Thr Thr Ile Tyr		
465	470	475
		480

Gln Ile Glu His Tyr Ser Met Asp Asp Leu Ala Ser Leu Lys Ile Pro
 485 490 495
 Glu Gln Phe Arg His Ala Ile Trp Lys Gly Ile Leu Asp His Arg Gln
 500 505 510
 Leu His Glu Phe Ser Ser Pro Ser His Leu Leu Arg Thr Pro Ser Ser
 515 520 525
 Ala Ser Thr Val Ser Val Gly Ser Ser Glu Thr Arg Gly Glu Arg Val
 530 535 540
 Ile Asp Ala Val Arg Phe Thr Leu Arg Gln Thr Ile Ser Phe Pro Pro
 545 550 555 560
 Arg Asp Glu Trp Asn Asp Phe Asn Phe Asp Met Asp Ala Arg Arg Asn
 565 570 575
 Lys Gln Gln Arg Ile Lys Glu Glu Gly Glu
 580 585
 <210> 339
 <211> 641
 <212> PRT
 <213> Homo sapiens
 <400> 339
 Met Ser Gln Ser Thr Gln Thr Asn Glu Phe Leu Ser Pro Glu Val Phe
 5 10 15
 Gln His Ile Trp Asp Phe Leu Glu Gln Pro Ile Cys Ser Val Gln Pro
 20 25 30
 Ile Asp Leu Asn Phe Val Asp Glu Pro Ser Glu Asp Gly Ala Thr Asn
 35 40 45
 Lys Ile Glu Ile Ser Met Asp Cys Ile Arg Met Gln Asp Ser Asp Leu
 50 55 60
 Ser Asp Pro Met Trp Pro Gln Tyr Thr Asn Leu Gly Leu Leu Asn Ser
 65 70 75 80
 Met Asp Gln Gln Ile Gln Asn Gly Ser Ser Ser Thr Ser Pro Tyr Asn
 85 90 95
 Thr Asp His Ala Gln Asn Ser Val Thr Ala Pro Ser Pro Tyr Ala Gln
 100 105 110
 Pro Ser Ser Thr Phe Asp Ala Leu Ser Pro Ser Pro Ala Ile Pro Ser
 115 120 125
 Asn Thr Asp Tyr Pro Gly Pro His Ser Phe Asp Val Ser Phe Gln Gln
 130 135 140

Ser Ser Thr Ala Lys Ser Ala Thr Trp Thr Tyr Ser Thr Glu Leu Lys
 145 150 155 160
 Lys Leu Tyr Cys Gln Ile Ala Lys Thr Cys Pro Ile Gln Ile Lys Val
 165 170 175
 Met Thr Pro Pro Pro Gln Gly Ala Val Ile Arg Ala Met Pro Val Tyr
 180 185 190
 Lys Lys Ala Glu His Val Thr Glu Val Val Lys Arg Cys Pro Asn His
 195 200 205
 Glu Leu Ser Arg Glu Phe Asn Glu Gly Gln Ile Ala Pro Pro Ser His
 210 215 220
 Leu Ile Arg Val Glu Gly Asn Ser His Ala Gln Tyr Val Glu Asp Pro
 225 230 235 240
 Ile Thr Gly Arg Gln Ser Val Leu Val Pro Tyr Glu Pro Pro Gln Val
 245 250 255
 Gly Thr Glu Phe Thr Thr Val Leu Tyr Asn Phe Met Cys Asn Ser Ser
 260 265 270
 Cys Val Gly Gly Met Asn Arg Arg Pro Ile Leu Ile Ile Val Thr Leu
 275 280 285
 Glu Thr Arg Asp Gly Gln Val Leu Gly Arg Arg Cys Phe Glu Ala Arg
 290 295 300
 Ile Cys Ala Cys Pro Gly Arg Asp Arg Lys Ala Asp Glu Asp Ser Ile
 305 310 315 320
 Arg Lys Gln Gln Val Ser Asp Ser Thr Lys Asn Gly Asp Gly Thr Lys
 325 330 335
 Arg Pro Phe Arg Gln Asn Thr His Gly Ile Gln Met Thr Ser Ile Lys
 340 345 350
 Lys Arg Arg Ser Pro Asp Asp Glu Leu Leu Tyr Leu Pro Val Arg Gly
 355 360 365
 Arg Glu Thr Tyr Glu Met Leu Leu Lys Ile Lys Glu Ser Leu Glu Leu
 370 375 380
 Met Gln Tyr Leu Pro Gln His Thr Ile Glu Thr Tyr Arg Gln Gln Gln
 385 390 395 400
 Gln Gln Gln His Gln His Leu Leu Gln Lys Gln Thr Ser Ile Gln Ser
 405 410 415
 Pro Ser Ser Tyr Gly Asn Ser Ser Pro Pro Leu Asn Lys Met Asn Ser
 420 425 430
 Met Asn Lys Leu Pro Ser Val Ser Gln Leu Ile Asn Pro Gln Gln Arg

155

435 440 445
 Asn Ala Leu Thr Pro Thr Thr Ile Pro Asp Gly Met Gly Ala Asn Ile
 450 455 460
 Pro Met Met Gly Thr His Met Pro Met Ala Gly Asp Met Asn Gly Leu
 465 470 475 480
 Ser Pro Thr Gln Ala Leu Pro Pro Pro Leu Ser Met Pro Ser Thr Ser
 485 490 495
 His Cys Thr Pro Pro Pro Tyr Pro Thr Asp Cys Ser Ile Val Gly
 500 505 510
 Phe Leu Ala Arg Leu Gly Cys Ser Ser Cys Leu Asp Tyr Phe Thr Thr
 515 520 525
 Gln Gly Leu Thr Thr Ile Tyr Gln Ile Glu His Tyr Ser Met Asp Asp
 530 535 540
 Leu Ala Ser Leu Lys Ile Pro Glu Gln Phe Arg His Ala Ile Trp Lys
 545 550 555 560
 Gly Ile Leu Asp His Arg Gln Leu His Glu Phe Ser Ser Pro Ser His
 565 570 575
 Leu Leu Arg Thr Pro Ser Ser Ala Ser Thr Val Ser Val Gly Ser Ser
 580 585 590
 Glu Thr Arg Gly Glu Arg Val Ile Asp Ala Val Arg Phe Thr Leu Arg
 595 600 605
 Gln Thr Ile Ser Phe Pro Pro Arg Asp Glu Trp Asn Asp Phe Asn Phe
 610 615 620
 Asp Met Asp Ala Arg Arg Asn Lys Gln Gln Arg Ile Lys Glu Glu Gly
 625 630 635 640
 Glu

<210> 340

<211> 448

<212> PRT

<213> Homo sapiens

<400> 340

Met Ser Gln Ser Thr Gln Thr Asn Glu Phe Leu Ser Pro Glu Val Phe
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Gln His Ile Trp Asp Phe Leu Glu Gln Pro Ile Cys Ser Val Gln Pro
 20 25 30

Ile Asp Leu Asn Phe Val Asp Glu Pro Ser Glu Asp Gly Ala Thr Asn
 35 40 45

Lys Ile Glu Ile Ser Met Asp Cys Ile Arg Met Gln Asp Ser Asp Leu
 50 55 60

Ser Asp Pro Met Trp Pro Gln Tyr Thr Asn Leu Gly Leu Leu Asn Ser
 65 70 75 80

Met Asp Gln Gln Ile Gln Asn Gly Ser Ser Ser Thr Ser Pro Tyr Asn
 85 90 95

Thr Asp His Ala Gln Asn Ser Val Thr Ala Pro Ser Pro Tyr Ala Gln
 100 105 110

Pro Ser Ser Thr Phe Asp Ala Leu Ser Pro Ser Pro Ala Ile Pro Ser
 115 120 125

Asn Thr Asp Tyr Pro Gly Pro His Ser Phe Asp Val Ser Phe Gln Gln
 130 135 140

Ser Ser Thr Ala Lys Ser Ala Thr Trp Thr Tyr Ser Thr Glu Leu Lys
 145 150 155 160

Lys Leu Tyr Cys Gln Ile Ala Lys Thr Cys Pro Ile Gln Ile Lys Val
 165 170 175

Met Thr Pro Pro Pro Gln Gly Ala Val Ile Arg Ala Met Pro Val Tyr
 180 185 190

Lys Lys Ala Glu His Val Thr Glu Val Val Lys Arg Cys Pro Asn His
 195 200 205

Glu Leu Ser Arg Glu Phe Asn Glu Gly Gln Ile Ala Pro Pro Ser His
 210 215 220

Leu Ile Arg Val Glu Gly Asn Ser His Ala Gln Tyr Val Glu Asp Pro
 225 230 235 240

Ile Thr Gly Arg Gln Ser Val Leu Val Pro Tyr Glu Pro Pro Gln Val
 245 250 255

Gly Thr Glu Phe Thr Thr Val Leu Tyr Asn Phe Met Cys Asn Ser Ser
 260 265 270

Cys Val Gly Gly Met Asn Arg Arg Pro Ile Leu Ile Ile Val Thr Leu
 275 280 285

Glu Thr Arg Asp Gly Gln Val Leu Gly Arg Arg Cys Phe Glu Ala Arg
 290 295 300

Ile Cys Ala Cys Pro Gly Arg Asp Arg Lys Ala Asp Glu Asp Ser Ile
 305 310 315 320

Arg Lys Gln Gln Val Ser Asp Ser Thr Lys Asn Gly Asp Gly Thr Lys
 325 330 335

Arg Pro Phe Arg Gln Asn Thr His Gly Ile Gln Met Thr Ser Ile Lys
340 345 350

Lys Arg Arg Ser Pro Asp Asp Glu Leu Leu Tyr Leu Pro Val Arg Gly
355 360 365

Arg Glu Thr Tyr Glu Met Leu Leu Lys Ile Lys Glu Ser Leu Glu Leu
370 375 380

Met Gln Tyr Leu Pro Gln His Thr Ile Glu Thr Tyr Arg Gln Gln Gln
385 390 395 400

Gln Gln Gln His Gln His Leu Leu Gln Lys His Leu Leu Ser Ala Cys
405 410 415

Phe Arg Asn Glu Leu Val Glu Pro Arg Arg Glu Thr Pro Lys Gln Ser
420 425 430

Asp Val Phe Phe Arg His Ser Lys Pro Pro Asn Arg Ser Val Tyr Pro
435 440 445

<210> 341

<211> 356

<212> PRT

<213> Homo sapiens

<400> 341

Met Leu Tyr Leu Glu Asn Asn Ala Gln Thr Gln Phe Ser Glu Pro Gln
5 10 15

Tyr Thr Asn Leu Gly Leu Leu Asn Ser Met Asp Gln Gln Ile Gln Asn
20 25 30

Gly Ser Ser Ser Thr Ser Pro Tyr Asn Thr Asp His Ala Gln Asn Ser
35 40 45

Val Thr Ala Pro Ser Pro Tyr Ala Gln Pro Ser Ser Thr Phe Asp Ala
50 55 60

Leu Ser Pro Ser Pro Ala Ile Pro Ser Asn Thr Asp Tyr Pro Gly Pro
65 70 75 80

His Ser Phe Asp Val Ser Phe Gln Gln Ser Ser Thr Ala Lys Ser Ala
85 90 95

Thr Trp Thr Tyr Ser Thr Glu Leu Lys Lys Leu Tyr Cys Gln Ile Ala
100 105 110

Lys Thr Cys Pro Ile Gln Ile Lys Val Met Thr Pro Pro Pro Gln Gly
115 120 125

Ala Val Ile Arg Ala Met Pro Val Tyr Lys Lys Ala Glu His Val Thr
130 135 140

Glu Val Val Lys Arg Cys Pro Asn His Glu Leu Ser Arg Glu Phe Asn

145 150 155 160
 Glu Gly Gln Ile Ala Pro Pro Ser His Leu Ile Arg Val Glu Gly Asn
 165 170 175
 Ser His Ala Gln Tyr Val Glu Asp Pro Ile Thr Gly Arg Gln Ser Val
 180 185 190
 Leu Val Pro Tyr Glu Pro Pro Gln Val Gly Thr Glu Phe Thr Thr Val
 195 200 205
 Leu Tyr Asn Phe Met Cys Asn Ser Ser Cys Val Gly Gly Met Asn Arg
 210 215 220
 Arg Pro Ile Leu Ile Ile Val Thr Leu Glu Thr Arg Asp Gly Gln Val
 225 230 235 240
 Leu Gly Arg Arg Cys Phe Glu Ala Arg Ile Cys Ala Cys Pro Gly Arg
 245 250 255
 Asp Arg Lys Ala Asp Glu Asp Ser Ile Arg Lys Gln Gln Val Ser Asp
 260 265 270
 Ser Thr Lys Asn Gly Asp Gly Thr Lys Arg Pro Ser Arg Gln Asn Thr
 275 280 285
 His Gly Ile Gln Met Thr Ser Ile Lys Lys Arg Arg Ser Pro Asp Asp
 290 295 300
 Glu Leu Leu Tyr Leu Pro Val Arg Gly Arg Glu Thr Tyr Glu Met Leu
 305 310 315 320
 Leu Lys Ile Lys Glu Ser Leu Glu Leu Met Gln Tyr Leu Pro Gln His
 325 330 335
 Thr Ile Glu Thr Tyr Arg Gln Gln Gln Gln Gln His Gln His Leu
 340 345 350
 Leu Gln Lys Gln
 355
 <210> 342
 <211> 680
 <212> PRT
 <213> Homo sapiens
 <400> 342
 Met Asn Phe Glu Thr Ser Arg Cys Ala Thr Leu Gln Tyr Cys Pro Asp
 5 10 15
 Pro Tyr Ile Gln Arg Phe Val Glu Thr Pro Ala His Phe Ser Trp Lys
 20 25 30
 Glu Ser Tyr Tyr Arg Ser Thr Met Ser Gln Ser Thr Gln Thr Asn Glu
 35 40 45

Phe Leu Ser Pro Glu Val Phe Gln His Ile Trp Asp Phe Leu Glu Gln
 50 55 60
 Pro Ile Cys Ser Val Gln Pro Ile Asp Leu Asn Phe Val Asp Glu Pro
 65 70 75 80
 Ser Glu Asp Gly Ala Thr Asn Lys Ile Glu Ile Ser Met Asp Cys Ile
 85 90 95
 Arg Met Gln Asp Ser Asp Leu Ser Asp Pro Met Trp Pro Gln Tyr Thr
 100 105 110
 Asn Leu Gly Leu Leu Asn Ser Met Asp Gln Gln Ile Gln Asn Gly Ser
 115 120 125
 Ser Ser Thr Ser Pro Tyr Asn Thr Asp His Ala Gln Asn Ser Val Thr
 130 135 140
 Ala Pro Ser Pro Tyr Ala Gln Pro Ser Ser Thr Phe Asp Ala Leu Ser
 145 150 155 160
 Pro Ser Pro Ala Ile Pro Ser Asn Thr Asp Tyr Pro Gly Pro His Ser
 165 170 175
 Phe Asp Val Ser Phe Gln Gln Ser Ser Thr Ala Lys Ser Ala Thr Trp
 180 185 190
 Thr Tyr Ser Thr Glu Leu Lys Lys Leu Tyr Cys Gln Ile Ala Lys Thr
 195 200 205
 Cys Pro Ile Gln Ile Lys Val Met Thr Pro Pro Pro Gln Gly Ala Val
 210 215 220
 Ile Arg Ala Met Pro Val Tyr Lys Lys Ala Glu His Val Thr Glu Val
 225 230 235 240
 Val Lys Arg Cys Pro Asn His Glu Leu Ser Arg Glu Phe Asn Glu Gly
 245 250 255
 Gln Ile Ala Pro Pro Ser His Leu Ile Arg Val Glu Gly Asn Ser His
 260 265 270
 Ala Gln Tyr Val Glu Asp Pro Ile Thr Gly Arg Gln Ser Val Leu Val
 275 280 285
 Pro Tyr Glu Pro Pro Gln Val Gly Thr Glu Phe Thr Thr Val Leu Tyr
 290 295 300
 Asn Phe Met Cys Asn Ser Ser Cys Val Gly Gly Met Asn Arg Arg Pro
 305 310 315 320
 Ile Leu Ile Ile Val Thr Leu Glu Thr Arg Asp Gly Gln Val Leu Gly
 325 330 335

Arg Arg Cys Phe Glu Ala Arg Ile Cys Ala Cys Pro Gly Arg Asp Arg
 340 345 350
 Lys Ala Asp Glu Asp Ser Ile Arg Lys Gln Gln Val Ser Asp Ser Thr
 355 360 365
 Lys Asn Gly Asp Gly Thr Lys Arg Pro Phe Arg Gln Asn Thr His Gly
 370 375 380
 Ile Gln Met Thr Ser Ile Lys Lys Arg Arg Ser Pro Asp Asp Glu Leu
 385 390 395 400
 Leu Tyr Leu Pro Val Arg Gly Arg Glu Thr Tyr Glu Met Leu Leu Lys
 405 410 415
 Ile Lys Glu Ser Leu Glu Leu Met Gln Tyr Leu Pro Gln His Thr Ile
 420 425 430
 Glu Thr Tyr Arg Gln Gln Gln Gln Gln His Gln His Leu Leu Gln
 435 440 445
 Lys Gln Thr Ser Ile Gln Ser Pro Ser Ser Tyr Gly Asn Ser Ser Pro
 450 455 460
 Pro Leu Asn Lys Met Asn Ser Met Asn Lys Leu Pro Ser Val Ser Gln
 465 470 475 480
 Leu Ile Asn Pro Gln Gln Arg Asn Ala Leu Thr Pro Thr Thr Ile Pro
 485 490 495
 Asp Gly Met Gly Ala Asn Ile Pro Met Met Gly Thr His Met Pro Met
 500 505 510
 Ala Gly Asp Met Asn Gly Leu Ser Pro Thr Gln Ala Leu Pro Pro Pro
 515 520 525
 Leu Ser Met Pro Ser Thr Ser Gln Cys Thr Pro Pro Pro Tyr Pro
 530 535 540
 Thr Asp Cys Ser Ile Val Ser Phe Leu Ala Arg Leu Gly Cys Ser Ser
 545 550 555 560
 Cys Leu Asp Tyr Phe Thr Thr Gln Gly Leu Thr Thr Ile Tyr Gln Ile
 565 570 575
 Glu His Tyr Ser Met Asp Asp Leu Ala Ser Leu Lys Ile Pro Glu Gln
 580 585 590
 Phe Arg His Ala Ile Trp Lys Gly Ile Leu Asp His Arg Gln Leu His
 595 600 605
 Glu Phe Ser Ser Pro Ser His Leu Leu Arg Thr Pro Ser Ser Ala Ser
 610 615 620
 Thr Val Ser Val Gly Ser Ser Glu Thr Arg Gly Glu Arg Val Ile Asp


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<210> 343
<211> 461
<212> PRT
<213> Homo sapiens
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<400> 343
Met Leu Tyr Leu Glu Asn Asn Ala Gln Thr Gln Phe Ser Glu Pro Gln
      5                      10                      15

Tyr Thr Asn Leu Gly Leu Leu Asn Ser Met Asp Gln Gln Ile Gln Asn
      20                      25                      30

Gly Ser Ser Ser Thr Ser Pro Tyr Asn Thr Asp His Ala Gln Asn Ser
      35                      40                      45

Val Thr Ala Pro Ser Pro Tyr Ala Gln Pro Ser Ser Thr Phe Asp Ala
      50                      55                      60

Leu Ser Pro Ser Pro Ala Ile Pro Ser Asn Thr Asp Tyr Pro Gly Pro
      65                      70                      75                      80

His Ser Phe Asp Val Ser Phe Gln Gln Ser Ser Thr Ala Lys Ser Ala
      85                      90                      95

Thr Trp Thr Tyr Ser Thr Glu Leu Lys Lys Leu Tyr Cys Gln Ile Ala
      100                      105                      110

Lys Thr Cys Pro Ile Gln Ile Lys Val Met Thr Pro Pro Pro Gln Gly
      115                      120                      125

Ala Val Ile Arg Ala Met Pro Val Tyr Lys Lys Ala Glu His Val Thr
      130                      135                      140

Glu Val Val Lys Arg Cys Pro Asn His Glu Leu Ser Arg Glu Phe Asn
      145                      150                      155                      160

Glu Gly Gln Ile Ala Pro Pro Ser His Leu Ile Arg Val Glu Gly Asn
      165                      170                      175

Ser His Ala Gln Tyr Val Glu Asp Pro Ile Thr Gly Arg Gln Ser Val
      180                      185                      190

Leu Val Pro Tyr Glu Pro Pro Gln Val Gly Thr Glu Phe Thr Thr Val
      195                      200                      205

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Leu Tyr Asn Phe Met Cys Asn Ser Ser Cys Val Gly Gly Met Asn Arg
 210 215 220
 Arg Pro Ile Leu Ile Ile Val Thr Leu Glu Thr Arg Asp Gly Gln Val
 225 230 235 240
 Leu Gly Arg Arg Cys Phe Glu Ala Arg Ile Cys Ala Cys Pro Gly Arg
 245 250 255
 Asp Arg Lys Ala Asp Glu Asp Ser Ile Arg Lys Gln Gln Val Ser Asp
 260 265 270
 Ser Thr Lys Asn Gly Asp Gly Thr Lys Arg Pro Phe Arg Gln Asn Thr
 275 280 285
 His Gly Ile Gln Met Thr Ser Ile Lys Lys Arg Arg Ser Pro Asp Asp
 290 295 300
 Glu Leu Leu Tyr Leu Pro Val Arg Gly Arg Glu Thr Tyr Glu Met Leu
 305 310 315 320
 Leu Lys Ile Lys Glu Ser Leu Glu Leu Met Gln Tyr Leu Pro Gln His
 325 330 335
 Thr Ile Glu Thr Tyr Arg Gln Gln Gln Gln Gln His Gln His Leu
 340 345 350
 Leu Gln Lys Gln Thr Ser Ile Gln Ser Pro Ser Ser Tyr Gly Asn Ser
 355 360 365
 Ser Pro Pro Leu Asn Lys Met Asn Ser Met Asn Lys Leu Pro Ser Val
 370 375 380
 Ser Gln Leu Ile Asn Pro Gln Gln Arg Asn Ala Leu Thr Pro Thr Thr
 385 390 395 400
 Ile Pro Asp Gly Met Gly Ala Asn Ile Pro Met Met Gly Thr His Met
 405 410 415
 Pro Met Ala Gly Asp Met Asn Gly Leu Ser Pro Thr Gln Ala Leu Pro
 420 425 430
 Pro Pro Leu Ser Met Pro Ser Thr Ser His Cys Thr Pro Pro Pro Pro
 435 440 445
 Tyr Pro Thr Asp Cys Ser Ile Val Arg Ile Trp Gln Val
 450 455 460

<210> 344

<211> 516

<212> PRT

<213> Homo sapiens

<400> 344

Met Ser Gln Ser Thr Gln Thr Asn Glu Phe Leu Ser Pro Glu Val Phe
 5 10 15
 Gln His Ile Trp Asp Phe Leu Glu Gln Pro Ile Cys Ser Val Gln Pro
 20 25 30
 Ile Asp Leu Asn Phe Val Asp Glu Pro Ser Glu Asp Gly Ala Thr Asn
 35 40 45
 Lys Ile Glu Ile Ser Met Asp Cys Ile Arg Met Gln Asp Ser Asp Leu
 50 55 60
 Ser Asp Pro Met Trp Pro Gln Tyr Thr Asn Leu Gly Leu Leu Asn Ser
 65 70 75 80
 Met Asp Gln Gln Ile Gln Asn Gly Ser Ser Ser Thr Ser Pro Tyr Asn
 85 90 95
 Thr Asp His Ala Gln Asn Ser Val Thr Ala Pro Ser Pro Tyr Ala Gln
 100 105 110
 Pro Ser Ser Thr Phe Asp Ala Leu Ser Pro Ser Pro Ala Ile Pro Ser
 115 120 125
 Asn Thr Asp Tyr Pro Gly Pro His Ser Phe Asp Val Ser Phe Gln Gln
 130 135 140
 Ser Ser Thr Ala Lys Ser Ala Thr Trp Thr Tyr Ser Thr Glu Leu Lys
 145 150 155 160
 Lys Leu Tyr Cys Gln Ile Ala Lys Thr Cys Pro Ile Gln Ile Lys Val
 165 170 175
 Met Thr Pro Pro Pro Gln Gly Ala Val Ile Arg Ala Met Pro Val Tyr
 180 185 190
 Lys Lys Ala Glu His Val Thr Glu Val Val Lys Arg Cys Pro Asn His
 195 200 205
 Glu Leu Ser Arg Glu Phe Asn Glu Gly Gln Ile Ala Pro Pro Ser His
 210 215 220
 Leu Ile Arg Val Glu Gly Asn Ser His Ala Gln Tyr Val Glu Asp Pro
 225 230 235 240
 Ile Thr Gly Arg Gln Ser Val Leu Val Pro Tyr Glu Pro Pro Gln Val
 245 250 255
 Gly Thr Glu Phe Thr Thr Val Leu Tyr Asn Phe Met Cys Asn Ser Ser
 260 265 270
 Cys Val Gly Gly Met Asn Arg Arg Pro Ile Leu Ile Ile Val Thr Leu
 275 280 285
 Glu Thr Arg Asp Gly Gln Val Leu Gly Arg Arg Cys Phe Glu Ala Arg

290 295 300
 Ile Cys Ala Cys Pro Gly Arg Asp Arg Lys Ala Asp Glu Asp Ser Ile
 305 310 315 320
 Arg Lys Gln Gln Val Ser Asp Ser Thr Lys Asn Gly Asp Gly Thr Lys
 325 330 335
 Arg Pro Phe Arg Gln Asn Thr His Gly Ile Gln Met Thr Ser Ile Lys
 340 345 350
 Lys Arg Arg Ser Pro Asp Asp Glu Leu Leu Tyr Leu Pro Val Arg Gly
 355 360 365
 Arg Glu Thr Tyr Glu Met Leu Leu Lys Ile Lys Glu Ser Leu Glu Leu
 370 375 380
 Met Gln Tyr Leu Pro Gln His Thr Ile Glu Thr Tyr Arg Gln Gln Gln
 385 390 395 400
 Gln Gln Gln His Gln His Leu Leu Gln Lys Gln Thr Ser Ile Gln Ser
 405 410 415
 Pro Ser Ser Tyr Gly Asn Ser Ser Pro Pro Leu Asn Lys Met Asn Ser
 420 425 430
 Met Asn Lys Leu Pro Ser Val Ser Gln Leu Ile Asn Pro Gln Gln Arg
 435 440 445
 Asn Ala Leu Thr Pro Thr Thr Ile Pro Asp Gly Met Gly Ala Asn Ile
 450 455 460
 Pro Met Met Gly Thr His Met Pro Met Ala Gly Asp Met Asn Gly Leu
 465 470 475 480
 Ser Pro Thr Gln Ala Leu Pro Pro Pro Leu Ser Met Pro Ser Thr Ser
 485 490 495
 His Cys Thr Pro Pro Pro Pro Tyr Pro Thr Asp Cys Ser Ile Val Arg
 500 505 510
 Ile Trp Gln Val
 515

<210> 345

<211> 1800

<212> DNA

<213> Homo sapiens

<400> 345

gcgcctcatt gccactgcag tgactaaagc tgggaagacg ctggtcagtt cacctgcccc 60
 actggttggt ttttaacaa attctgatac aggcgacatc ctactgacc gagcaaagat 120
 tgacattcgt atcatcactg tgcaccattg gcttctaggc actccagtgg ggtaggagaa 180

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ggaggtctga aaccctcgca gagggatctt gccctcattc tttgggtctg aaacactggc 240
agtcgttgga aacaggactc agggataaac cagcgcaatg gattggggga cgctgcacac 300
tttcatcggg ggtgtcaaca aacactccac cagcatcggg aagggtgga tcacagtcac 360
ctttattttc cgagtcacga tcttagtggt ggctgcccag gaagtgtggg gtgacgagca 420
agaggacttc gtctgcaaca cactgcaacc gggatgcaaa aatgtgtgct atgaccactt 480
tttcccgggtg tcccacatcc ggctgtgggc cctccagctg atcttctgtc ccacccagc 540
gctgctgggtg gccatgcatg tggcctacta caggcacgaa accactcgca agttcaggcg 600
aggagagaag aggaatgatt tcaaagacat agaggacatt aaaaagcaca aggttcggat 660
agaggggtcg ctgtggtgga cgtacaccag cagcatcttt ttccgaatca tctttgaagc 720
agcctttatg tatgtgtttt acttccttta caatgggtac cacctgccct ggggtgtgaa 780
atgtgggatt gacccctgcc ccaaccttgt tgactgcttt atttctaggc caacagagaa 840
gaccgtgttt accattttta tgatttctgc gtctgtgatt tgcagtctgc ttaacgtggc 900
agagttgtgc tacctgtctg tgaaagtgtg ttttaggaga tcaaagagag cacagacgca 960
aaaaaatcac cccaatcatg ccctaaagga gagtaagcag aatgaaatga atgagctgat 1020
ttcagatagt ggtcaaaatg caatcacagg tttcccaagc taaacatttc aaggtaaaat 1080
gtagctgcgt cataaggaga cttctgtctt ctccagaagg caataccaac ctgaaagttc 1140
cttctgtagc ctgaagagtt tgtaaatgac tttcataata aatagacact tgagttaact 1200
ttttgtagga tacttgctcc attcatacac aacgtaatca aatatgtggt ccatctctga 1260
aaacaagaga ctgcttgaca aaggagcatt gcagtcactt tgacagggtc cttttaagtg 1320
gactctctga caaagtgggt actttctgaa aatttatata actgttgttg ataaggaaca 1380
tttatccagg aattgatacg tttattagga aaagatatat ttataggctt ggatgttttt 1440
agttccgact ttgaatttat ataaagtatt tttataatga ctggtcttcc ttacctggaa 1500
aaacatgcga tgtagttttt agaattacac cacaagtatc taaatttcca acttacaag 1560
ggtcctatct tgtaaatatt gttttgcatt gtctgttggc aaatttgtga actgtcatga 1620
tacgcttaag gtgggaaagt gttcattgca caatatatit ttactgcttt ctgaatgtag 1680
acggaacagt gtggaagcag aaggcttttt taactcatcc gtttgccga tcgttgacaga 1740
ccactgggag atgtggatgt ggttgctccc ttttgcctgt ccccggtggt taacctttct 1800

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<210> 346

<211> 261

<212> PRT

<213> Homo sapiens

<400> 346

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Met Asp Trp Gly Thr Leu His Thr Phe Ile Gly Gly Val Asn Lys His
      5                      10                      15

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Ser Thr Ser Ile Gly Lys Val Trp Ile Thr Val Ile Phe Ile Phe Arg
      20                      25                      30

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Val Met Ile Leu Val Val Ala Ala Gln Glu Val Trp Gly Asp Glu Gln
      35                      40                      45

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Glu Asp Phe Val Cys Asn Thr Leu Gln Pro Gly Cys Lys Asn Val Cys
      50                      55                      60

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Tyr Asp His Phe Phe Pro Val Ser His Ile Arg Leu Trp Ala Leu Gln
      65                      70                      75                      80

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Leu Ile Phe Val Ser Thr Pro Ala Leu Leu Val Ala Met His Val Ala
      85                      90                      95

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Tyr Tyr Arg His Glu Thr Thr Arg Lys Phe Arg Arg Gly Glu Lys Arg
      100                      105                      110

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Asn Asp Phe Lys Asp Ile Glu Asp Ile Lys Lys His Lys Val Arg Ile
 115 120 125
 Glu Gly Ser Leu Trp Trp Thr Tyr Thr Ser Ser Ile Phe Phe Arg Ile
 130 135 140
 Ile Phe Glu Ala Ala Phe Met Tyr Val Phe Tyr Phe Leu Tyr Asn Gly
 145 150 155 160
 Tyr His Leu Pro Trp Val Leu Lys Cys Gly Ile Asp Pro Cys Pro Asn
 165 170 175
 Leu Val Asp Cys Phe Ile Ser Arg Pro Thr Glu Lys Thr Val Phe Thr
 180 185 190
 Ile Phe Met Ile Ser Ala Ser Val Ile Cys Met Leu Leu Asn Val Ala
 195 200 205
 Glu Leu Cys Tyr Leu Leu Leu Lys Val Cys Phe Arg Arg Ser Lys Arg
 210 215 220
 Ala Gln Thr Gln Lys Asn His Pro Asn His Ala Leu Lys Glu Ser Lys
 225 230 235 240
 Gln Asn Glu Met Asn Glu Leu Ile Ser Asp Ser Gly Gln Asn Ala Ile
 245 250 255
 Thr Gly Phe Pro Ser
 260

<210> 347

<211> 1740

<212> DNA

<213> Homo sapiens

<400> 347

atgaacaaac tgtatatcgg aaacctcagc gagaacgccg cccctcggga cctagaaagt 60
 atcttcaagg acgccaagat cccggtgtcg ggacccttcc tgggtaagac tggctacgcg 120
 ttcgtggact gcccgacga gagctgggccc ctcaaggcca tcgaggcgct ttcaggtaaa 180
 atagaactgc acgggaaaacc catagaagtt gagcactcgg tcccaaaaag gcaaaggatt 240
 cggaacttc agatacgaat tatccgcct catttacagt gggaggtgct ggatagttta 300
 ctagtccagt atggagtggg ggagagctgt gagcaagtga aactgactc ggaaactgca 360
 gttgtaaatg taacctattc cagtaaggac caagctagac aagcactaga caaactgaat 420
 ggatttcagt tagagaattt caccttgaat gtagcctata tccctgatga aacggccgcc 480
 cagcaaaacc ccttcagca gcccgagggt cgcggggggc ttgggcagag gggctcctca 540
 aggcaggggt ctccaggatc cgtatccaag cagaaaccat gtgatttgcc tctgcgcctg 600
 ctggttccca cccaatttgt tggagccatc ataggaaaag aagggtgccac cattcggaac 660
 atcaccaaac agaccagtc taaaatcgat gtccaccgta aagaaaatgc gggggctgct 720
 gagaagtoga ttactatcct ctctactcct gaaggcacct ctgcggcttg taagtctatt 780
 ctggagatta tgcataagga agctcaagat ataaaattca cagaagagat ccccttgaag 840
 attttagctc ataataactt tggttgacgt cttattggta aagaaggag aaatcttaaa 900
 aaaattgagc aagacacaga cactaaaatc acgatatctc cattgcagga attgacgctg 960

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180	185	190
Pro Cys Asp Leu Pro Leu Arg Leu Leu Val Pro Thr Gln Phe Val Gly		
195	200	205
Ala Ile Ile Gly Lys Glu Gly Ala Thr Ile Arg Asn Ile Thr Lys Gln		
210	215	220
Thr Gln Ser Lys Ile Asp Val His Arg Lys Glu Asn Ala Gly Ala Ala		
225	230	235
Glu Lys Ser Ile Thr Ile Leu Ser Thr Pro Glu Gly Thr Ser Ala Ala		
245	250	255
Cys Lys Ser Ile Leu Glu Ile Met His Lys Glu Ala Gln Asp Ile Lys		
260	265	270
Phe Thr Glu Glu Ile Pro Leu Lys Ile Leu Ala His Asn Asn Phe Val		
275	280	285
Gly Arg Leu Ile Gly Lys Glu Gly Arg Asn Leu Lys Lys Ile Glu Gln		
290	295	300
Asp Thr Asp Thr Lys Ile Thr Ile Ser Pro Leu Gln Glu Leu Thr Leu		
305	310	315
Tyr Asn Pro Glu Arg Thr Ile Thr Val Lys Gly Asn Val Glu Thr Cys		
325	330	335
Ala Lys Ala Glu Glu Glu Ile Met Lys Lys Ile Arg Glu Ser Tyr Glu		
340	345	350
Asn Asp Ile Ala Ser Met Asn Leu Gln Ala His Leu Ile Pro Gly Leu		
355	360	365
Asn Leu Asn Ala Leu Gly Leu Phe Pro Pro Thr Ser Gly Met Pro Pro		
370	375	380
Pro Thr Ser Gly Pro Pro Ser Ala Met Thr Pro Pro Tyr Pro Gln Phe		
385	390	395
Glu Gln Ser Glu Thr Glu Thr Val His Leu Phe Ile Pro Ala Leu Ser		
405	410	415
Val Gly Ala Ile Ile Gly Lys Gln Gly Gln His Ile Lys Gln Leu Ser		
420	425	430
Arg Phe Ala Gly Ala Ser Ile Lys Ile Ala Pro Ala Glu Ala Pro Asp		
435	440	445
Ala Lys Val Arg Met Val Ile Ile Thr Gly Pro Pro Glu Ala Gln Phe		
450	455	460
Lys Ala Gln Gly Arg Ile Tyr Gly Lys Ile Lys Glu Glu Asn Phe Val		
465	470	475
		480

Ser Pro Lys Glu Glu Val Lys Leu Glu Ala His Ile Arg Val Pro Ser
 485 490 495

Phe Ala Ala Gly Arg Val Ile Gly Lys Gly Gly Lys Thr Val Asn Glu
 500 505 510

Leu Gln Asn Leu Ser Ser Ala Glu Val Val Val Pro Arg Asp Gln Thr
 515 520 525

Pro Asp Glu Asn Asp Gln Val Val Val Lys Ile Thr Gly His Phe Tyr
 530 535 540

Ala Cys Gln Val Ala Gln Arg Lys Ile Gln Glu Ile Leu Thr Gln Val
 545 550 555 560

Lys Gln His Gln Gln Gln Lys Ala Leu Gln Ser Gly Pro Pro Gln Ser
 565 570 575

Arg Arg Lys

<210> 349

<211> 207

<212> DNA

<213> Homo sapiens

<400> 349

atgtggcagc cctcttcttt caagtggctc ttgtcctggt gccctgggag ttctcaaatt 60
 gctgcagcag cctccaccca gcctgaggat gacatcaata cacagaggaa gaagagtcag 120
 gaaaagatga gagaagttac agactctcct gggcgacccc gagagcttac cattcctcag 180
 acttcttcac atggtgctaa cagattt 207

<210> 350

<211> 69

<212> PRT

<213> Homo sapiens

<400> 350

Met Trp Gln Pro Leu Phe Phe Lys Trp Leu Leu Ser Cys Cys Pro Gly
 5 10 15

Ser Ser Gln Ile Ala Ala Ala Ala Ser Thr Gln Pro Glu Asp Asp Ile
 20 25 30

Asn Thr Gln Arg Lys Lys Ser Gln Glu Lys Met Arg Glu Val Thr Asp
 35 40 45

Ser Pro Gly Arg Pro Arg Glu Leu Thr Ile Pro Gln Thr Ser Ser His
 50 55 60

Gly Ala Asn Arg Phe
 65

